Logging in to Dialog Trying 31060000009999...Open DIALOG INFORMATION SERVICES PLEASE LOGON: ****** ENTER PASSWORD: Welcome to DIALOG Dialog level 05.31.00D Last logoff: 19apr11 11:43:04 Logon file405 19apr11 12:34:18 DETAIL set on HILIGHT set on as '****' COST = SHORT. MEDIOBAB is set ON as an alias for 155, 347, 144, 35, 5, 74, 71, 357, 6, 351, 24, 136, 399, 315, 358, 73, 34, 434 FISH is set ON as an alias for 10, 143, 203, 50, 28, 35, 351, 24, 136, 44, NUTRACEUT is set ON as an alias for 79, 164, 91, 53, 51, 351, 399, 467,149 MEDBIOFT is set ON as an alias for 349, 444, 457 SYSTEM: HOME Cost is in DialUnits Menu System II: D2 version 1.8.0 term=ASCII *** DIALOG HOMEBASE(SM) Main Menu *** Information: 1. Announcements (new files, reloads, etc.) 2. Database, Rates, & Command Descriptions 3. Help in Choosing Databases for Your Topic 4. Customer Services (telephone assistance, training, seminars, etc.) 5. Product Descriptions Connections: 6. DIALOG(R) Document Delivery Data Star(R) (c) 2003 Dialog, a Thomson business. All rights reserved. /H = Help /L = Logoff /NOMENU = Command Mode Enter an option number to view information or to connect to an online service. Enter a BEGIN command plus a file number to search a database

(e.g., B1 for ERIC). ? b mediobab

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358 does not exist
>>>1 of the specified files is not available
      19apr11 12:34:25 User226352 Session D1308.1
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    $0.05 TELNET
    $0.05 Estimated cost this search
    $0.05 Estimated total session cost 0.263 DialUnits
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SYSTEM: OS - DIALOG OneSearch
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         (c) format only 2011 Dialog
*File 155: Medline has been reloaded with the 2011 MeSH
thesaurus.
 File 347: JAPIO Dec 1976-2010/Dec (Updated 110323)
         (c) 2011 JPO & JAPIO
  File 144:Pascal 1973-2011/Apr W2
         (c) 2011 INIST/CNRS
       35:Dissertation Abs Online 1861-2011/Mar
         (c) 2011 ProQuest Info&Learning
  File
        5:Biosis Previews(R) 1926-2011/Apr W2
         (c) 2011 The Thomson Corporation
  File 74: Int. Pharm. Abs 1970-2011/Apr B2
        (c) 2011 The Thomson Corporation
 File 71:ELSEVIER BIOBASE 1994-2011/Apr W3
         (c) 2011 Elsevier B.V.
  File 357: Derwent Biotech Res. _1982-2011/Nov W4
         (c) 2011 Thomson Reuters
*File 357: This file will no longer be produced after 12/31/2010. For
more information, see HELP NEWS 357.
       6:NTIS 1964-2011/Apr W4
         (c) 2011 NTIS, Intl Cpyrght All Rights Res
  File 351:Derwent WPI 1963-2011/UD=201125
         (c) 2011 Thomson Reuters
       24:CSA Life Sciences Abstracts 1966-2011/Mar
         (c) 2011 CSA.
  File 136:BioEngineering Abstracts 1966-2007/Jan
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*File 136: This file is closed.
  File 399:CA SEARCH(R) 1967-2010/UD=15417
         (c) 2011 American Chemical Society
*File 399: Use is subject to the terms of your user/customer agreement.
IPCR/8 classification codes now searchable as IC=. See HELP NEWSIPCR.
 File 315: ChemEng & Biotec Abs 1970-2011/May
         (c) 2011 DECHEMA
*File 315: December 2007 - the reloaded database is now online. Please
consult the updated Bluesheet for details on new and changed fields.
  File 73:EMBASE 1974-2011/Apr 19
         (c) 2011 Elsevier B.V.
*File 73: The 2011 Thesaurus has been installed with UD20110407.
 File 34:SciSearch(R) Cited Ref Sci 1990-2011/Apr W2
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  File 434:SciSearch(R) Cited Ref Sci 1974-1989/Dec
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347: JAPIO_Dec 1976-2010/Dec(Updated 110323)
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144: Pascal_1973-2011/Apr W2

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35: Dissertation Abs Online_1861-2011/Mar

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5: Biosis Previews(R)_1926-2011/Apr W2 260 W135

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74: Int.Pharm.Abs 1970-2011/Apr B2

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71: ELSEVIER BIOBASE_1994-2011/Apr W3 155 W135

- 11125 CONJUGATE
- 2333 MENTINGOCOCCAL
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357: Derwent Biotech Res.__1982-2011/Nov W4

- 27 W135 218 MENINGOCOCCAL
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- 73: EMBASE_1974-2011/Apr 19 385 W135
 - 37047 CONJUGATE
 - 8430 MENTINGOCOCCAL

CONJUGATE

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434: SciSearch(R) Cited Ref Sci 1974-1989/Dec
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74: Int.Pharm.Abs_1970-2011/Apr B2
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E5	351		2	AU=COSTANTINO, PAOLO, CHIRON S.P.A., VIA
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E6	351		2	AU=COSTANTINO, PAOLO, CHIRON S.R.L., VIA
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E7	351		1	AU=COSTANTINO, PAOLO, CHIRON SRL
E8	351		1	AU=COSTANTINO, PAOLO, COLLE VAL DIELSA, IT
E9	351			AU=COSTANTINO, PAOLO, IT
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E12	351		1	AU=COSTANTINO, PAOLO, NOVARTIS VACCINES, AND
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155: MEDLINE(R)_1950-2011/Apr 15

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347: JAPIO Dec 1976-2010/Dec(Updated 110323)

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144: Pascal 1973-2011/Apr W2

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^{74:} Int.Pharm.Abs_1970-2011/Apr B2

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71: ELSEVIER BIOBASE_1994-2011/Apr W3

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357: Derwent Biotech Res.__1982-2011/Nov W4

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351: Derwent WPI_1963-2011/UD=201125

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24: CSA Life Sciences Abstracts_1966-2011/Mar

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136: BioEngineering Abstracts_1966-2007/Jan

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315: ChemEng & Biotec Abs_1970-2011/May

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73: EMBASE_1974-2011/Apr 19

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155: MEDLINE(R)_1950-2011/Apr 15

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35: Dissertation Abs Online_1861-2011/Mar

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0021027583

WPI ACC NO: 2010-M58148/201066

Immunogenic composition for raising an immune response in a mammal comprises meningococcal lipooligosaccharide (LOS) and a pneumococcal serotype 14 ****capsular**** saccharide (CS14)

Patent Assignee: NOVARTIS AG (NOVS)

Inventor: COSTANTINO P

Patent Family (2 patents, 113 countries)
Patent Application

Priority Applications (no., kind, date): US 2009162996 P 20090324

Patent Details

Number Kind Lan Pg Dwg Filing Notes

47

WO 2010109325 A2 EN

National Designated States, Confirmed: AE AG AL AM AO AT AU AZ BA BB BG BH BR BW BY BZ CA CH CL CN CO CR CU CZ DE DR DM DO DZ EC EE BG ES FI GB GB GE GB GH GM GT HN HR HU ID IL IN IS JP KE KG KM KN KP KR KZ LA LC LK LS LT LU LY MA MD ME MG MK MN MM XM YM ZN AN GN IN ON ZO MP EP GP PH LP TRORS RU SC SD SE SG SK SL SM ST SV SY TH TJ TM TN TR TT TZ UA UG US UZ VC VN ZA ZM ZW

WO 2010109325 A3 EN

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Alerting Abstract WO A2

NOVELTY - Immunogenic composition comprises a meningococcal lippoligosaccharide (LOS) and a pneumococcal serotype 14 ***capsular**** saccharide (CS14), where the LOS and/or the CS14 do(es) not include a Gal-betai-4 N-acetylglucosamine (GlcNAc)-betai-3Gal-betai-4Glc tetrasaccharide.

DESCRIPTION - INDEPENDENT CLAIMS are:

1.an unadjuvanted immunogenic composition comprising LOS and a CS14, where the LOS and the CS14 both include a

Gal-betal-4GlcNAc-betal-3Gal-betal-4Glc tetrasaccharide; and

2.a method for raising an immune response in a mammal comprising administering a composition to the mammal.

ACTIVITY - Immunostimulant; Antibacterial; Neuroprotective; Immunosuppressive. No biological data given.

MECHANISM OF ACTION - Vaccine.

USE - The immunogenic composition is useful for raising an immune response in a mammal (claimed) and for treating and preventing diseases caused by "***Neisseria**** meningitides ~ and/or ~Streptococcus pneumoniae ~ e.g. bacterial (or, more specifically, meningococcal and/or meningitis, or septicemia).

ADVANTAGE - The composition provides improved combination vaccines for protecting against both meningococcus serogroup B and pneumococcus.

Technology Focus

BIOTECHNOLOGY - Preferred Composition: In the composition, the

tetrasaccharide is present in CS14. It comprises an adjuvant. The LOS is prepared from a meningococcal strain lacking LqtB enzyme activity. The LOS is prepared from a meningococcal strain lacking GalE enzyme activity. The LOS is prepared from a meningococcal strain lacking LgtA and/or LgtE enzyme activity. The LOS lacks the terminal galactose of the Gal-betal-4GlcNAc-betal-3Gal-betal-4Glc tetrasaccharide. The LOS is present within meningococcal outer membrane vesicles. The LOS is conjugated to proteins in the vesicles. The vesicles are prepared from a meningococcus that over-expresses TbpA. The LOS is conjugated to a carrier protein. The conjugation may be via a lipid A portion in the LOS or by a 3-deoxy-d-manno-octulosonic acid (KDO) residue. Alternatively, the immunogenic composition comprises one or more meningococcal polypeptide(s) and a CS14, where (a) the CS14 includes a Gal-betal-4GlcNAc1-3Gal-betal-4Glc tetrasaccharide, (b) the meningococcal polypeptide can elicit an immune response that is effective against serogroup B meningococcus, and (c) the composition does not include a meningococcal lipooligosaccharide. The meningococcal polypeptide(s) comprise a factor H binding protein (fHBP). The fHBP comprises an amino acid sequence having at least 85% sequence identity to SEQ ID NO. 1 (not defined) and/or comprising an amino acid sequence consisting of a fragment of at least 7 contiquous amino acids from SEQ ID NO. 1. The fHBP comprises an amino acid sequence having at least 85% sequence identity to SEO ID NO. 2 (not defined) and/or comprising an amino acid sequence consisting of a fragment of at least 7 contiguous amino acids from SEO ID NO. 2. The fHBP comprises an amino acid sequence having at least 85% sequence identity to SEQ ID NO. 3 (not defined) and/or comprising an amino acid sequence consisting of a fragment of at least 7 contiguous amino acids from SEQ ID NO. 3. The fHBP is lipidated. The fHBP elicits antibodies which can bind to a meningococcal polypeptide consisting of amino acid sequence SEQ ID NO. 1, 2 or 3. It also comprises a LOS and a CS14, where (a) the LOS and the CS14 both include a Gal-betal-4GlcNAc-betal-3Gal-betal-4Glc tetrasaccharide, (b) the concentration of LOS is less than mu g/ml, and (c) the concentration of CS14 is less than 5 mu g/ml. The CS14 is conjugated to a carrier protein. The carrier protein is CRM197, tetanus toxoid, diphtheria toxoid or ~Haemophilus influenzae ~ protein D. It also comprises a LOS and a pneumococcal polypeptide antigen, where (a) the LOS includes a Gal-betal-4GlcNAc-betal-3Gal-betal-4Glc tetrasaccharide, (b) the pneumococcal polypeptide can elicit an immune response that is effective against serotype 14 pneumococcus, and (c) the composition does not include a CS14. It comprises an aluminum salt adjuvant. The aluminum salt is an aluminum phosphate.

Title Terms/Index Terms/Additional Words: IMMUNOGENIC; COMPOSITION; RAISE; IMMUNE; RESPOND; MAMMAL, COMPRISE; MENINGOCOCCUS; PNEUMOCOCCUS; SEROLOGICAL; CAPSULE; SACCHARIDE

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Class Codes
IPC + Level Value Position Status Version
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A61K-0039/09 A I L B 20060101
A61K-0039/09 A I L B 20060101
A61K-0039/095 A I L B 20060101
C07K-0014/315 A I L B 20060101
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ECLA: A61K-039/095, C07K-014/315B
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File Segment: CPI

DWPI Class: B04; D16

Manual Codes (CPI/A-M): B04-C02F; B04-E99; B04-N03; B05-B02A3; B14-A01A5; B14-A01B2; B14-C03; B14-G01; B14-N16; B14-S06; B14-S11B1; D05-H07

Publication No. WO 2010109325 A2 (Update 201066 B)

WIPO

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Publication Date: 20100930
**COMBINATIONS INCLUDING PNEUMOCOCCAL SEROTYPE 14 SACCHARIDE
  COMBINAISONS COMPRENANT UN SACCHARIDE DE PNEUMOCOOUE SEROTYPE 14**
Assignee: ~(except US)~ NOVARTIS AG, Lightstrasse 35, CH-4056 Basel, CH
    Residence: CH Nationality: CH (NOVS)
  ~(only US)~ COSTANTINO, Paolo, Novartis Vaccines, Via Fiorentina, 1,
    I-53100 Siena, IT Residence: IT Nationality: IT
Inventor: COSTANTINO, Paolo, Novartis Vaccines, Via Fiorentina, 1, I-53100
    Siena, IT Residence: IT Nationality: IT
Agent: MARSHALL, Cameron, John et al., Carpmaels Ransford, One Southampton
    Row, London WC1B 5HA, GB
Language: EN (47 pages, 4 drawings)
Application: WO 2010IB735 A 20100324 (Local application)
Priority: US 2009162996 P 20090324
Designated States: (National Confirmed) AE AG AL AM AO AT AU AZ BA BB BG BH
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    GE GH GM GT HN HR HU ID IL IN IS JP KE KG KM KN KP KR KZ LA LC LK LR LS
    LT LU LY MA MD ME MG MK MN MW MX MY MZ NA NG NI NO NZ OM PE PG PH PL PT
    RO RS RU SC SD SE SG SK SL SM ST SV SY TH TJ TM TN TR TT TZ UA UG US UZ
    VC VN ZA ZM ZW
Original IPC: A61K-39/02(B,I,H,EP,20060101,A,F)
    A61K-39/02(B, I, M, 98, 20060101, C)
Current IPC: A61K-39/02(B.I.H.EP.20060101,A.F)
   A61K-39/02(B, I, M, 98, 20060101, C)
Current ECLA class: A61K-39/095 C07K-14/315B
Original Abstract: Meningococcal lipooligosaccharide and pneumococcal
    serotype 14 capsular saccharide share an antigen that can cross-react
    with human tissue. The invention provides various ways of minimising
    the production of autoreactive antibodies when these two antigens are
    co-administered.
   Le lipo-oligo-saccharide de meningocoque et le saccharide capsulaire de
   pneumocoque serotype 14 partagent un antigene qui peut presenter une
    reaction croisee avec un tissu humain. L'invention concerne divers
   movens de reduire la production d'anticorps autoreactifs lorsque ces
    deux antigenes sont co-administres.
Publication No. WO 2010109325 A3 (Update 201107 E)
Publication Date: 20110120
**COMBINATIONS INCLUDING PNEUMOCOCCAL SEROTYPE 14 SACCHARIDE**
Assignee: NOVARTIS AG; CH (NOVS)
Inventor: COSTANTINO P, IT
Language: EN
Application: WO 2010IB735 A 20100324 (Local application)
Priority: US 2009162996 P 20090324
Designated States: (National Confirmed) AE AG AL AM AO AT AU AZ BA BB BG BH
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    RO RS RU SC SD SE SG SK SL SM ST SV SY TH TJ TM TN TR TT TZ UA UG US UZ
    VC VN ZA ZM ZW
Original IPC: A61K-39/02(B,I,H,EP,20060101,A,F)
    A61K-39/09(B,I,H,EP,20060101,A,L) A61K-39/095(B,I,H,EP,20060101,A,L)
    A61K-39/295(B,I,H,EP,20060101,A,L) C07K-14/315(B,I,H,EP,20060101,A,L)
Current IPC: A61K-39/02(B,I,H,EP,20060101,A,F)
```

A61K-39/09(B,I,H,EP,20060101,A,L) A61K-39/095(B,I,H,EP,20060101,A,L) A61K-39/295(B,I,H,EP,20060101,A,L) C07K-14/315(B,I,H,EP,20060101,A,L)

Original Abstract: Meningococcal lipooligosaccharide and pneumococcal serotype 14 capsular saccharide share an antigen that can cross-react with human tissue. The invention provides various ways of minimising the production of autoreactive antibodies when these two antigens are co-administered.

10/7/2 (Item 2 from file: 351) DIALOG(R)File 351:Derwent WPI (c) 2011 Thomson Reuters. All rts. reserv. 0020394151 WPI ACC NO: 2010-E89462/201032 Purification of Streptococcus pyogenes carbohydrate useful for the preparation of vaccines involves use of anionic exchange chromatography Patent Assignee: NOVARTIS AG (NOVS) Inventor: BERTI F; COSTANTINO P; KABANOVA A; ROMANO M R Patent Family (1 patents, 124 countries) Pat.ent. Application Date Number Kind Number Kind Date WO 2010049806 A1 20100506 WO 2009IB7346 A 20091027 201032 B Priority Applications (no., kind, date): US 2008108763 P 20081027 Patent Details Pg Dwg Filing Notes Number Kind Lan WO 2010049806 A1 EN 56 1.5 National Designated States, Confirmed: AE AG AL AM AO AT AU AZ BA BB BG BH BR BW BY BZ CA CH CL CN CO CR CU CZ DE DK DM DO DZ EC EE EG ES FI GB GD GE GH GM GT HN HR HU ID IL IN IS JP KE KG KM KN KP KR KZ LA LC LK LR LS LT LU LY MA MD ME MG MK MN MW MX MY MZ NA NG NI NO NZ OM PE PG PH PL PT RO RS RU SC SD SE SG SK SL SM ST SV SY TJ TM TN TR TT TZ UA UG US UZ VC VN ZA ZM ZW Regional Designated States, Confirmed: AT BE BG CH CY CZ DE DK EE ES FI FR

Alerting Abstract WO A1

NOVELTY - Purification (m1) of ~Streptococcus pyogenes ~ (group A ~Streptococcus ~ (GAS)) carbohydrate involves use of anionic exchange chromatography.

BW GH GM KE LS MW MZ NA SD SL SZ TZ UG ZM ZW EA

GB GR HR HU IE IS IT LT LU LV MC MK MT NL NO PL PT RO SE SI SK SM TR OA

ACTIVITY - None given.

MECHANISM OF ACTION - Vaccines. Test details are described, but no results given.

USE - For purifying "Streptococcus pyogenes " carbohydrate (claimed) which are used in the preparation of vaccines, and as an antigen for in vitro diagnostic assays in e.g. immunization.

ADVANTAGE - The anionic exchange chromatography provides a good yield (i.e. greater than 70, preferably greater than 75, greater than 80, greater than 80, of GAS carbohydrate; and particularly pure GAS carbohydrate preparation. The GAS carbohydrate is often contaminated with hyaluronic acid, which is derived from the GAS ****capsular****
****polysaccharide****, thus anionic exchange chromatography is particularly effective at reducing hyaluronic acid contamination of GAS carbohydrate. This is particularly advantageous when the GAS carbohydrate is intended for use in a vaccine because hyaluronic acid is known to be immunogenic in its own right and induces antibodies that are cross-reactive with human tissue, so its presence in pharmaceutical products is detrimental to health. Anionic exchange chromatography is also particularly effective at reducing protein and nucleic acid contamination of GAS carbohydrate. The purification of GAS carbohydrate is performed under conditions that allow flow through of the saccharide during anionic

exchange chromatography, where impurities bind to the anion exchange matrix while GAS carbohydrate flows straight through the system into the eluant. The use of these conditions simplifies the purification process, as there is no need to use a mobile phase buffer of increasing joinic strength or increasing pH to elute the GAS carbohydrate from the matrix. The method provides a composition comprising a level of hyaluronic acid contamination that is less than 200 (preferably less than 150, more preferably less than 80, most preferably less than 40, especially 20, and particularly less than 10 ng/mi or less than 1 wt.% of hyaluronic acid relative to the weight of GAS carbohydrate; a level of polyrhamnose contamination that is less than 20 wt.% of polyrhamose relative to the weight of GAS carbohydrate; a level of protein contamination that is around 2 wt.% of protein relative to the weight of GAS carbohydrate; and a level of nucleic acid contamination that is less than 1 wt.% of nucleic acid relative to the weight of GAS carbohydrate; and level of nucleic acid contamination that is less than 1 wt.% of nucleic acid relative to the weight of GAS carbohydrate; and level of relative to the weight of GAS carbohydrate; and level of relative to the weight of GAS carbohydrate; and level of nucleic acid contamination that is less than 1 wt.% of nucleic acid relative to the weight of GAS carbohydrate.

Technology Focus

ORGANIC CHEMISTRY - Preferred Method: The suspension is prepared by treating ~S. pyogenes ~ such that the GAS carbohydrate is released. The GAS carbohydrate is released by reductive acid treatment. The method involves filtration step(s) prior to the anionic exchange chromatography step. The filtration is by orthogonal filtration using 0.65 mu m filter. The method involves at least one ultrafiltration step prior to the anionic exchange chromatography step. The ultrafiltration is by tangential flow filtration using 30 kDa cut-off membrane. The anionic exchange chromatography step is carried out using a Q-resin as anionic exchange matrix. The anionic exchange chromatography step is carried out using anionic exchange matrix resin (1 ml) for every 1 mg of GAS carbohydrate. The anionic exchange chromatography step is performed under conditions that allow flow through of the GAS carbohydrate. The mobile phase buffer for the anionic exchange chromatography comprises alcohol (preferably ethanol) in an amount of 15-25%. The method involves at least one gel filtration step after the anionic exchange chromatography step. The gel filtration step(s) are carried out using a dextran gel as gel filtration matrix (1 ml for every 0.2 mg of GAS carbohydrate); and are performed using the same mobile phase buffer as the anionic exchange chromatography step. The method involves concentrating the GAS carbohydrate after the anionic exchange chromatography step. The concentration step(s) are carried out by tangential flow filtration using a 5 or 10 kDa cut-off membrane. The method involves conjugating the purified GAS carbohydrate to a carrier molecule.

POLYMERS - Preferred Components: The purified GAS carbohydrate has a molecular weight of 10 kDa. The saccharide is partially or fully de-N-acetylated. The starting material is an aqueous suspension of the GAS carbohydrate, further comprising hyaluronic acid and/or polyrhamnose.

Title Terms/Index Terms/Additional Words: PURIFICATION, STREPTOCOCCUS; PYOGENES; CARBOHYDRATE; USEFUL; PREPARATION; VACCINE; ANION; EXCHANGE; CHROMATOGRAPHY

Class Codes International Classification (+ Attributes) IPC + Level Value Position Status Version A61K-003/90 A I L B 20060101 B01D-0015/36 A I F B 20060101 B01D-0061/14 A I L B 20060101 C07K-0001/18 A I L B 20060101 C07K-0001/34 A I L B 20060101 C12P-0019/00 A I L B 20060101 A61K-0039/09 C I L B 20100101 B01D-0051/26 C I F B 20100101 B01D-0061/14 C I L B 20100101 B01D-0061/14 C I L B 20100101

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C12P-0019/00 C I L B 20100101
ECLA: A61K-039/09A
ICO: K61K-039:555A, K61K-039:60P10, L01D-061:14D
File Segment: CPI; EPI
DWPI Class: B04; D16; J01; S03
Manual Codes (EPI/S-X): S03-E09C5
Manual Codes (CPI/A-M): B04-C02F; B11-B03; B12-K04A; B14-S11; D05-H13;
 J01-C; J01-D01A; J01-D04; J01-D07
Original Publication Data by Authority
WIPO
Publication No. WO 2010049806 A1 (Update 201032 B)
Publication Date: 20100506
**PURIFICATION METHOD
  PROCEDE DE PURIFICATION**
Assignee: ~(except US)~ NOVARTIS AG, Lichstrasse 35, CH-4056 Basel, CH
    Residence: CH Nationality: CH (NOVS)
  ~(only US)~ COSTANTINO, Paolo, Novartis Vaccines, Via Fiorentina, 1,
    I-53100 Siena, IT Residence: IT Nationality: IT
  ~(only US)~ BERTI, Francesco, Novartis Vaccines, Via Fiorentina, 1,
    I-53100 Siena, IT Residence: IT Nationality: IT
  ~(only US)~ KABANOVA, Anna, Novartis Vaccines, Via Fiorentina, 1, I-53100
    Siena, IT Residence: IT Nationality: RU
  ~(only US)~ ROMANO, Maria, Rosaria, Novartis Vaccines, Via Fiorentina, 1,
    I-53100 Siena, IT Residence: IT Nationality: IT
Inventor: BERTI, Francesco, Novartis Vaccines, Via Fiorentina, 1, I-53100
    Siena, IT Residence: IT Nationality: IT
  COSTANTINO, Paolo, Novartis Vaccines, Via Fiorentina, 1, I-53100 Siena,
    IT Residence: IT Nationality: IT
  KABANOVA, Anna, Novartis Vaccines, Via Fiorentina, 1, I-53100 Siena, IT
    Residence: IT Nationality: RU
  ROMANO, Maria, Rosaria, Novartis Vaccines, Via Fiorentina, 1, I-53100
    Siena, IT Residence: IT Nationality: IT
Agent: MARSHALL, Cameron, John et al., Carpmaels Ransford, 43-45
    Bloomsbury Square, London, WC1A 2RA, GB
Language: EN (56 pages, 15 drawings)
Application: WO 2009IB7346 A 20091027 (Local application)
Priority: US 2008108763 P 20081027
Designated States: (National Confirmed) AE AG AL AM AO AT AU AZ BA BB BG BH
    BR BW BY BZ CA CH CL CN CO CR CU CZ DE DK DM DO DZ EC EE EG ES FI GB GD
    GE GH GM GT HN HR HU ID IL IN IS JP KE KG KM KN KP KR KZ LA LC LK LR LS
    LT LU LY MA MD ME MG MK MN MW MX MY MZ NA NG NI NO NZ OM PE PG PH PL PT
    RO RS RU SC SD SE SG SK SL SM ST SV SY TJ TM TN TR TT TZ UA UG US UZ VC
    VN ZA ZM ZW
  (Regional Confirmed) AT BE BG CH CY CZ DE DK EE ES FI FR GB GR HR HU IE
    IS IT LT LU LV MC MK MT NL NO PL PT RO SE SI SK SM TR OA BW GH GM KE LS
    MW MZ NA SD SL SZ TZ UG ZM ZW EA
Original IPC: A61K-39/09(B,I,H,EP,20060101,A,L)
    A61K-39/09(B, I, M, 98, 20060101, C) B01D-15/26(B, I, M, 98, 20060101, C)
    B01D-15/36(B,I,H,EP,20060101,A,F) B01D-61/14(B,I,H,EP,20060101,A,L)
    B01D-61/14(B,I,M,98,20060101,C) C07K-1/00(B,I,M,98,20060101,C)
    C07K-1/18(B,I,H,EP,20060101,A,L) C07K-1/34(B,I,H,EP,20060101,A,L)
    C12P-19/00(B,I,H,EP,20060101,A,L) C12P-19/00(B,I,M,98,20060101,C)
Current IPC: A61K-39/09(B,I,H,EP,20060101,20100506,A,L)
    A61K-39/09(B,I,H,EP,20100101,20100506,C,L)
    B01D-15/26(B,I,H,EP,20100101,20100506,C,F)
    B01D-15/36(B, I, H, EP, 20060101, 20100506, A, F)
    B01D-61/14(B, I, H, EP, 20060101, 20100506, A, L)
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Current ECLA ICO class: K61K-39:555A K61K-39:60P10 L01D-61:14D
Original Abstract: A process for purifying a Streptococcus pyogenes GAS
   carbohydrate comprising a step of anionic exchange chromatography. The
   process provides a good yield of GAS carbohydrate. The saccharides of
   the invention have low levels of hyaluronic acid, protein and nucleic
   acid contamination.
  La presente invention concerne un procede permettant la purification du
   glucide Streptococcus pyogenes (GAS), comprenant une etape de
   chromatographie par echange anionique. Le procede permet d'obtenir un
   bon rendement de glucide GAS. Les saccharides selon l'invention
   presentent de faibles niveaux de contamination par l'acide
   hyaluronique, par des proteines et par l'acide nucleique.
10/7/3
           (Item 3 from file: 351)
DIALOG(R)File 351:Derwent WPI
(c) 2011 Thomson Reuters. All rts. reserv.
0019961320 - Drawing available
WPI ACC NO: 2009-S40218/201008
Production of conjugate of ****capsular**** saccharide of Salmonella typhi
e.g. for raising immune response, by combining linker, carbodiimide and
carrier protein, removing excess linker, and reacting with product of
saccharide with carbodiimide
Patent Assignee: NOVARTIS AG (NOVS)
Inventor: BERTI F; COSTANTINO P; MICOLI F
Patent Family (5 patents, 125 countries)
Patent
                              Application
Number
               Kind Date
                              Number
                                            Kind Date
                                                           Update
WO 2009150543
               A2 20091217 WO 2009IB6285
                                            A 20090612 201008 B
WO 2009150543
               A3 20100506 WO 2009IB6285
                                            A 20090612 201030 E
AU 2009259017
               A1 20091217 AU 2009259017
                                            A 20090612
                                                           201109 E
CA 2727565
               A1 20091217 CA 2727565
                                             A 20090612
                                                           201120 E
                              WO 2009IB6285
                                             A 20090612
                              CA 2727565
                                             A 20101210
EP 2303333
                A2 20110406 EP 2009762071
                                             A 20090612 201124 E
                              WO 2009IB6285
                                              A 20090612
Priority Applications (no., kind, date): GB 200810894 A 20080613
Patent Details
Number
              Kind Lan
                          Pa Dwa Filina Notes
WO 2009150543
                             18
               A2 EN
                         42
National Designated States, Confirmed: AE AG AL AM AO AT AU AZ BA BB BG BH
  BR BW BY BZ CA CH CL CN CO CR CU CZ DE DK DM DO DZ EC EE EG ES FI GB GD
  GE GH GM GT HN HR HU ID IL IN IS JP KE KG KM KN KP KR KZ LA LC LK LR LS
  LT LU LY MA MD ME MG MK MN MW MX MY MZ NA NG NI NO NZ OM PE PG PH PL PT
  RO RS RU SC SD SE SG SK SL SM ST SV SY TJ TM TN TR TT TZ UA UG US UZ VC
  VN ZA ZM ZW
Regional Designated States, Confirmed: AT BE BG CH CY CZ DE DK EE ES FI FR
  GB GR HR HU IE IS IT LT LU LV MC MK MT NL NO PL PT RO SE SI SK TR OA BW
  GH GM KE LS MW MZ NA SD SL SZ TZ UG ZM ZW EA
WO 2009150543
               A3 EN
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National Designated States, Confirmed: AE AG AL AM AO AT AU AZ BA BB BG BH BR BW BY BZ CA CH CL CN CO CR CU CZ DE DK DM DO DZ EC EE EG ES FI GB GD

B01D-61/14 (B, I, H, EP, 20100101, 20100506, C, L) C07K-1/00 (B, I, H, EP, 20100101, 20100506, C, L) C07K-1/18 (B, I, H, EP, 20060101, 20100506, A, L) C07K-1/34 (B, I, H, EP, 20060101, 20100506, A, L) C12P-19/00 (B, I, H, EP, 20060101, 20100506, A, L) C12P-19/00 (B, I, H, EP, 20100101, 20100506, C, L)

Current ECLA class: A61K-39/09A

GE GH GM GT HN HH HU ID IL IN IS JP KE KG KM KN KP KR KZ LA LC LK LR LE LI LU LY MA MD ME MG MG KM MM KM KY MY ZN AN RO NI NO NZ OM PE PG PH PL PT RO RS RU SC SD SE SG SK SL SM ST SV SY TJ TM TN TR TT TZ UA UG US UZ VC VN ZA ZM C X

WO 2009150543

Regional Designated States, Confirmed: AT BE BG CH CY CZ DE DK EE ES FI FR
GB GR HR HU IE IS IT LT LU LV MC MK MT NL NO PL PT RO SE SI SK TR OA BW
GH GM KE LS MW MZ NA SD SL SZ TZ UG ZM ZW EA

GH GM KE LS MW MZ NA SD SL SZ TZ UG ZM ZW EA
AU 2009259017 A1 EN Based on OPI patent

CA 2727565 A1 EN PCT Application WO 2009IB6285 PCT national entry CA 2727565 Based on OPI patent WO 2009150543

EP 2303333 A2 EN PCT Application WO 2009IB6285
Based on OPI patent WO 2009I50543

Regional Designated States, Confirmed: AT BE BG CH CY CZ DE DK EE ES FI FR GB GR HR HU IE IS IT LI LT LU LV MC MK MT NL NO PL PT RO SE SI SK TR AL BA RS

Alerting Abstract WO A2

NOVELTY - Production of conjugate of ****capsular**** saccharide of *Salmonella typhi ~ (Vi) comprises simultaneously combining a linker, carbodiimide and carrier protein; removing any excess linker from the product of first step; reacting Vi with carbodiimide; and reacting the product of second step with the product of third step.

DESCRIPTION - INDEPENDENT CLAIMS are included for:

- a conjugate comprising Vi linked to cross reacting material 197 (CRM197);
- 2.a pharmaceutical composition comprising the conjugate in combination with a carrier; and
- 3.a method for derivatizing a Vi saccharide comprising reacting the saccharide with a carbodimide at Vi:carbodimide molar ratio of greater than 3:1.

ACTIVITY - Immunostimulant; Antipyretic; Antibacterial. MECHANISM OF ACTION - Vaccine.

USE - The method is for production of conjugate of Vi for raising an immune response in a mammal (claimed) or for use in medicine for raising an antibody response in a mammal, and for manufacture of medicament for preventing or treating typhoid fever in a mammal. The mammal is preferably a human. Balb/c female mice were divided in 14 groups of eight mice each and were subcutaneously immunized with 2.5 mu g of Vi, Vi-conjugate or physical mixture of Vi and ADH-derivatized carrier protein. Only groups 13 and 14 received 10 mu g of immunization dose. Three injections of 200 mu l each were given every two weeks, with bleedings two weeks after each immunization. Groups 9-12 received alum as adjuvant, while the adjuvant for groups 13 and 14 was complete Freund's adjuvant (first injection) and incomplete Freund's adjuvant (second and third injections). An anti-Vi antibody values for groups 1-14 were respectively (T14;T28;T42): (3.3;-1.03; - 1.54), (- 3.4; - 1.94; - 0.04), (5.0; - 0.14; 0.03), (1.1; 2.51; 1.31), (53.7; 242.20; 191.53), (95.4; 245.39; 225.66), (75.5; 170.20; 160.49), (58.2;126.23;126.20), (56.4;162.52;98.58), (58.7;98.92;93.22), (44.6;191.81;151.56), (68.1;202.99;180.09), (65.3;271.48;264.39), and (134.5:202.51:234.78).

DESCRIPTION OF DRAWINGS - The drawing shows a graph of size-exclusion chromatography analysis of Vi-TT(ADH).

Technology Focus

BIOTECHNOLOGY - Preferred Component: The carrier protein is CRM197 or tetanus toxoid (TT).

INORGANIC CHEMISTRY - Preferred Component: The composition comprises

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saline and optionally adjuvant.
 ORGANIC CHEMISTRY - Preferred Component: The carbodiimide is
1-ethy1-3(3-dimethylaminopropy1)carbodiimide. The Vi:carbodiimide molar
ratio is greater than 5:1 (preferably greater than 9:1). Preferred Process:
The excess linker is removed by dialysis. The Vi is linked to CRM197 by an
adipic acid dihydrazide (ADH) as linker.
Title Terms/Index Terms/Additional Words: PRODUCE; CONJUGATE; CAPSULE;
  SACCHARIDE; SALMONELLA; TYPHI; RAISE; IMMUNE; RESPOND; COMBINATION; LINK;
  CARBODIIMIDE; CARRY; PROTEIN; REMOVE; EXCESS; REACT; PRODUCT
Class Codes
International Classification (+ Attributes)
IPC + Level Value Position Status Version
 A61K-0047/48 A I F B 20060101
  A61P-0043/00 A N L B 20060101
 A61P-0043/00 A I L B 20060101
 A61K-0047/48 C I
                        B 20060101
 A61K-0047/48 C I F B 20100101
  A61P-0043/00 C N
                        B 20060101
  A61P-0043/00 C N L B 20100101
ECLA: A61K-047/48R2D
File Segment: CPI
DWPI Class: B04; B07
Manual Codes (CPI/A-M): B04-C02F; B04-N03; B11-A01A; B14-A01A8; B14-C04;
 B14-G01; B14-S11B1
Original Publication Data by Authority
Australia
Publication No. AU 2009259017 A1 (Update 201109 E)
Publication Date: 20091217
Assignee: NOVARTIS AG (NOVS)
Inventor: COSTANTINO P
 MICOLI F
 BERTI F
Language: EN
Application: AU 2009259017 A 20090612 (Local application)
Priority: GB 200810894 A 20080613
Related Publication: WO 2009150543 A (Based on OPI patent )
Original IPC: A61K-47/48(B,I,H,EP,20060101,20091217,A,F)
    A61P-43/00(B, N, H, EP, 20060101, 20091217, A, L)
Current IPC: A61K-47/48(B,I,H,EP,20060101,20091217,A,F)
    A61P-43/00(B, N, H, EP, 20060101, 20091217, A, L)
Current ECLA class: A61K-47/48R2D
Canada
Publication No. CA 2727565 A1 (Update 201120 E)
Publication Date: 20091217
Assignee: NOVARTIS AG; CH (NOVS)
Inventor: MICOLI F, IT
 COSTANTINO P. IT
  BERTI F, IT
Language: EN
Application: CA 2727565 A 20090612 (Local application)
  WO 2009IB6285 A 20090612 (PCT Application)
 CA 2727565 A 20101210 (PCT national entry)
Priority: GB 200810894 A 20080613
Related Publication: WO 2009150543 A (Based on OPI patent )
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Original IPC: A61K-47/48(B,I,H,EP,20060101,20110129,A,F)
    A61P-43/00 (B, N, H, EP, 20060101, 20110129, A, L)
Current IPC: A61K-47/48(B,I,H,EP,20060101,20110129,A,F)
    A61P-43/00(B, N, H, EP, 20060101, 20110129, A, L)
EP0
Publication No. EP 2303333 A2 (Update 201124 E)
Publication Date: 20110406
**CONJUGATED VI SACCHARIDES**
Assignee: NOVARTIS AG: CH (NOVS)
Inventor: MICOLI F, IT
  COSTANTINO P, IT
  BERTI F, IT
Language: EN
Application: EP 2009762071 A 20090612 (Local application)
  WO 2009IB6285 A 20090612 (PCT Application)
Priority: GB 200810894 A 20080613
Related Publication: WO 2009150543 A (Based on OPI patent )
Designated States: (Regional Confirmed) AT BE BG CH CY CZ DE DK EE ES FI FR
    GB GR HR HU IE IS IT LI LT LU LV MC MK MT NL NO PL PT RO SE SI SK TR AL
    BA RS
Original IPC: A61K-47/48(B,I,H,EP,20060101,20100106,A,F)
    A61P-43/00(B, I, H, EP, 20060101, 20100106, A, L)
Current IPC: A61K-47/48(B.I.H.EP.20060101.20100106.A.F)
    A61P-43/00(B, I, H, EP, 20060101, 20100106, A, L)
WIPO
Publication No. WO 2009150543 A2 (Update 201008 B)
Publication Date: 20091217
**CONJUGATED VI SACCHARIDES
 CONJUGUES VI DE SACCHARIDES**
Assignee: ~(except US)~ NOVARTIS AG, Lichstrasse 35, CH-4056 Basel, CH
    Residence: CH Nationality: CH (NOVS)
  ~(only US)~ MICOLI, Francesca, Novartis Vaccines, Via Fiorentina 1,
    I-53100 Siena, IT Residence: IT Nationality: IT
  ~(only US)~ COSTANTINO, Paolo, Novartis Vaccines, Via Fiorentina 1,
    I-53100 Siena, IT Residence: IT Nationality: IT
  ~(only US)~ BERTI, Francesco, Novartis Vaccines, Via Fiorentina 1,
    I-53100 Siena, IT Residence: IT Nationality: IT
Inventor: BERTI, Francesco, Novartis Vaccines, Via Fiorentina 1, I-53100
    Siena, IT Residence: IT Nationality: IT
  COSTANTINO, Paolo, Novartis Vaccines, Via Fiorentina 1, I-53100 Siena, IT
   Residence: IT Nationality: IT
  MICOLI, Francesca, Novartis Vaccines, Via Fiorentina 1, I-53100 Siena, IT
    Residence: IT Nationality: IT
Agent: MARSHALL, Cameron, John, Carpmaels Ransford, 43-45 Bloomsbury
    Square, London WC1A 2RA, GB
Language: EN (42 pages, 18 drawings)
Application: WO 2009IB6285 A 20090612 (Local application)
Priority: GB 200810894 A 20080613
Designated States: (National Confirmed) AE AG AL AM AO AT AU AZ BA BB BG BH
    BR BW BY BZ CA CH CL CN CO CR CU CZ DE DK DM DO DZ EC EE EG ES FI GB GD
    GE GH GM GT HN HR HU ID IL IN IS JP KE KG KM KN KP KR KZ LA LC LK LR LS
    LT LU LY MA MD ME MG MK MN MW MX MY MZ NA NG NI NO NZ OM PE PG PH PL PT
    RO RS RU SC SD SE SG SK SL SM ST SV SY TJ TM TN TR TT TZ UA UG US UZ VC
    VN ZA ZM ZW
  (Regional Confirmed) AT BE BG CH CY CZ DE DK EE ES FI FR GB GR HR HU IE
    IS IT LT LU LV MC MK MT NL NO PL PT RO SE SI SK TR OA BW GH GM KE LS MW
   MZ NA SD SL SZ TZ UG ZM ZW EA
Original IPC: A61K-47/48(B,I,H,EP,20060101,A,F)
    A61K-47/48(B, I, M, 98, 20060101, C) A61P-43/00(B, N, H, EP, 20060101, A, L)
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A61P-43/00(B,N,M,98,20060101,C)

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Current IPC: A61K-47/48(B,I,H,EP,20060101,A,F)
    A61K-47/48(B, I, M, 98, 20060101, C) A61P-43/00(B, N, H, EP, 20060101, A, L)
    A61P-43/00(B,N,M,98,20060101,C)
Current ECLA class: A61K-47/48R2D
Original Abstract: Two Vi conjugates have been prepared by
    carbodiimide-mediated synthesis, using adipic acid dihydrazide
    derivatized CRM197 (a non-toxic variant of diphtheria toxin) and
    tetanus toxoid, as carrier proteins.
   Deux conjugues Vi ont ete prepares par une synthese induite par les
    carbodiimides, utilisant la CRM197 derivatisee de dihydrazide d'acide
    adipique (un variant non toxique de la toxine de diphterie) et le
    toxoide du tetanos, comme proteines-supports.
Publication No. WO 2009150543 A3 (Update 201030 E)
Publication Date: 20100506
**CONJUGATED VI SACCHARIDES**
Assignee: NOVARTIS AG; CH (NOVS)
Inventor: BERTI F, IT
 COSTANTINO P, IT
 MICOLI F, IT
Language: EN
Application: WO 2009IB6285 A 20090612 (Local application)
Priority: GB 200810894 A 20080613
Designated States: (National Confirmed) AE AG AL AM AO AT AU AZ BA BB BG BH
    BR BW BY BZ CA CH CL CN CO CR CU CZ DE DK DM DO DZ EC EE EG ES FI GB GD
    GE GH GM GT HN HR HU ID IL IN IS JP KE KG KM KN KP KR KZ LA LC LK LR LS
    LT LU LY MA MD ME MG MK MN MW MX MY MZ NA NG NI NO NZ OM PE PG PH PL PT
    RO RS RU SC SD SE SG SK SL SM ST SV SY TJ TM TN TR TT TZ UA UG US UZ VC
    VN ZA ZM ZW
  (Regional Confirmed) AT BE BG CH CY CZ DE DK EE ES FI FR GB GR HR HU IE
    IS IT LT LU LV MC MK MT NL NO PL PT RO SE SI SK TR OA BW GH GM KE LS MW
    MZ NA SD SL SZ TZ UG ZM ZW EA
Original IPC: A61K-47/48(B,I,H,EP,20060101,A,F)
    A61K-47/48(B,I,M,98,20060101,C) A61P-43/00(B,N,H,EP,20060101,A,L)
    A61P-43/00(B,N,M,98,20060101,C)
Current IPC: A61K-47/48(B,I,H,EP,20060101,20100506,A,F)
    A61K-47/48(B, I, H, EP, 20100101, 20100506, C, F)
    A61P-43/00 (B, N, H, EP, 20060101, 20100506, A, L)
    A61P-43/00(B, N, H, EP, 20100101, 20100506, C, L)
Current ECLA class: A61K-47/48R2D
Original Abstract: Two Vi conjugates have been prepared by
    carbodiimide-mediated synthesis, using adipic acid dihydrazide
    derivatized CRM197 (a non-toxic variant of diphtheria toxin) and
   tetanus toxoid, as carrier proteins.
            (Item 4 from file: 351)
 10/7/4
DIALOG(R)File 351:Derwent WPI
(c) 2011 Thomson Reuters. All rts. reserv.
0019961318 - Drawing available
WPI ACC NO: 2009-S40212/201008
Quantification of ****capsular**** saccharide of Salmonella typhi present
in sample, comprises de-0-acetylating any ****capsular**** saccharide of
Salmonella typhi present in sample, and obtaining nuclear magnetic
resonance spectrum of sample
Patent Assignee: NOVARTIS AG
                              (NOVS)
Inventor: BERTI F; MICOLI F; PROIETTI D
Patent Family (6 patents, 125 countries)
Patent
                               Application
               Kind Date
Number
                               Number
                                              Kind Date
                                                             Update
WO 2009150533 A2 20091217 WO 2009IB6087 A 20090612 201008 B
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WO 2009150533
             A3 20100204 WO 2009IB6087 A 20090612 201010 E
AU 2009259007
             A1 20091217 AU 2009259007 A 20090612 201111 E
                                        A 20090612 201120 E
CA 2727563
              A1 20091217 CA 2727563
                          WO 2009IB6087 A 20090612
                                        A 20101210
                          CA 2727563
IN 201004854
              P2
                  20110311 WO 2009IB6087
                                        A 20090612 201123 E
                           IN 2010KN4854
                                        A 20101220
EP 2300817
              A2 20110330 EP 2009762061
                                         A 20090612 201124 E
                          WO 2009IB6087
                                         A 20090612
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Priority Applications (no., kind, date): IT 2008MI1079 A 20080613

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Patent Details
Number
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Kind Lan Pg Dwg Filing Notes WO 2009150533 A2 EN 23 13

A3 EN

P2 EN

National Designated States, Confirmed: AE AG AL AM AO AT AU AZ BA BB BG BH BR BW BY BZ CA CH CL CN CO CR CU CZ DE DK DM DO DZ EC EE EG ES FI GB GD GE GH GM GT HN HR HU ID IL IN IS JP KE KG KM KN KP KR KZ LA LC LK LR LS LT LU LY MA MD ME MG MK MN MW MX MY MZ NA NG NI NO NZ OM PE PG PH PL PT RO RS RU SC SD SE SG SK SL SM ST SV SY TJ TM TN TR TT TZ UA UG US UZ VC

VN ZA ZM ZW Regional Designated States, Confirmed: AT BE BG CH CY CZ DE DK EE ES FI FR GB GR HR HU IE IS IT LT LU LV MC MK MT NL NO PL PT RO SE SI SK TR OA BW

GH GM KE LS MW MZ NA SD SL SZ TZ UG ZM ZW EA WO 2009150533

IN 201004854

EP 2300817

National Designated States, Confirmed: AE AG AL AM AO AT AU AZ BA BB BG BH BR BW BY BZ CA CH CL CN CO CR CU CZ DE DK DM DO DZ EC EE EG ES FI GB GD GE GH GM GT HN HR HU ID IL IN IS JP KE KG KM KN KP KR KZ LA LC LK LR LS LT LU LY MA MD ME MG MK MN MW MX MY MZ NA NG NI NO NZ OM PE PG PH PL PT RO RS RU SC SD SE SG SK SL SM ST SV SY TJ TM TN TR TT TZ UA UG US UZ VC VN ZA ZM ZW

Regional Designated States, Confirmed: AT BE BG CH CY CZ DE DK EE ES FI FR GB GR HR HU IE IS IT LT LU LV MC MK MT NL NO PL PT RO SE SI SK TR OA BW GH GM KE LS MW MZ NA SD SL SZ TZ UG ZM ZW EA

AU 2009259007 A1 EN CA 2727563 A1 EN

Based on OPI patent WO 2009150533 PCT Application WO 2009IB6087 PCT national entry CA 2727563 Based on OPI patent WO 2009150533 PCT Application WO 2009IB6087 PCT Application WO 2009IB6087

Based on OPI patent WO 2009150533 Regional Designated States, Confirmed: AT BE BG CH CY CZ DE DK EE ES FI FR GB GR HR HU IE IS IT LI LT LU LV MC MK MT NL NO PL PT RO SE SI SK TR AL BA RS

Alerting Abstract WO A2

NOVELTY - Quantification of ****capsular**** saccharide of ~Salmonella typhi ~ (Vi) present in a sample comprises de-0-acetylating any Vi saccharide present in the sample, and obtaining a nuclear magnetic resonance (NMR) spectrum of the sample.

DESCRIPTION - An INDEPENDENT CLAIM is included for a method for quantifying Vi saccharide present in a sample by liquid chromatography.

USE - Quantification of Vi saccharide present in a sample (claimed).

ADVANTAGE - The two methods are simple and accurate for quantifying Vi saccharide and its conjugates that allow the detection of very low (<= 5 mu q/ml, including as low as 1 mu q/ml) Vi concentrations. The liquid chromatography is an improvement over conventional techniques, such as acridine orange method, as it permits the quantification of Vi in a sample containing proteins, reagents and other contaminants.

DESCRIPTION OF DRAWINGS - The drawing shows a graph of HPAEC-PAD analysis of Vi saccharide after de-O-acetylation and hydrolysis with 4 M TFA at 120 degrees Celsius for 2 hours.

Technology Focus BIOTECHNOLOGY - Preferred Component: The Vi saccharide is from ~S. typhi ~ or ~Citrobacter freundii ~ . INORGANIC CHEMISTRY - Preferred Component: The Vi saccharide is

de-O-acetylated by sodium deuteroxide.

INSTRUMENTATION AND TESTING - Preferred Process: The method further comprises using NMR spectrum to calculate the amount of Vi saccharide present in the sample, and adding a known amount of reference compound to the sample. An N-acetyl resonance is used to calculate the amount of Vi saccharide present in the sample. The NMR spectroscopy is hydrogen-1 NMR spectroscopy. The liquid chromatography is high performance anion exchange chromatography (HPAEC), and the method uses pulsed amperometric detection (PAD) or (HPAEC-PAD). The method comprises hydrolyzing Vi saccharide present in the sample, de-acetylating Vi saccharide present in the sample, and analyzing the sample by liquid chromatography. The method further comprises a second hydrolysis step that follows the de-acetylation step. The hydrolysis step(s) is carried out by treatment with 4 M trifluoroacetic acid (TFA) at 120(deg) C for 2 hours. The de-acetylation step is carried out by treatment with 2 M sodium hydroxide at 110 (deg) C for 6 hours. Hydrolysis and de-acetylation involves treatment with sodium hydroxide at 100-150(deg) C for 2-6 hours.

ORGANIC CHEMISTRY - Preferred Component: The reference compound is citric acid or ethanol.

Title Terms/Index Terms/Additional Words: QUANTIFICATION; CAPSULE; SACCHARIDE; SALMONELLA; TYPHI; PRESENT; SAMPLE; COMPRISE; DE; ACETYLATE; OBTAIN; NUCLEAR; MAGNETIC; RESONANCE; SPECTRUM

Class Codes

International Classification (Main): G01N-033/15 International Classification (+ Attributes) IPC + Level Value Position Status Version G01N-0033/15 A I F B 20060101

ECLA: G01R-033/46

ICO: S01N-224:352, S01N-333:255, S01N-400:10, S01R-330:400A, S01R-330:404B

File Segment: CPI; EPI DWPI Class: B04; J04; S03

Manual Codes (EPI/S-X): S03-E07C; S03-E14A1

Manual Codes (CPI/A-M): B04-C02F; B11-C08A; B11-C08D2; B12-K04; J04-B01A; J04-B01C; J04-C01; J04-C03

Original Publication Data by Authority

Australia

Publication No. AU 2009259007 A1 (Update 201111 E) Publication Date: 20091217

Assignee: NOVARTIS AG (NOVS) Inventor: MICOLI F

BERTI F PROIETTI D Language: EN

Application: AU 2009259007 A 20090612 (Local application)

Priority: IT 2008MI1079 A 20080613 Related Publication: WO 2009150533 A (Based on OPI patent)

Original IPC: G01N-33/15(B,I,H,EP,20060101,20100204,A,F) Current IPC: G01N-33/15(B,I,H,EP,20060101,20100204,A,F)

Current ECLA class: G01R-33/46

Current ECLA ICO class: S01N-224:352 S01N-333:255 S01N-400:10 S01R-330:400A

```
Canada
Publication No. CA 2727563 A1 (Update 201120 E)
Publication Date: 20091217
Assignee: NOVARTIS AG; CH (NOVS)
Inventor: BERTI F, IT
 MICOLI F, IT
 PROIETTI D, II
Language: EN
Application: CA 2727563 A 20090612 (Local application)
  WO 2009IB6087 A 20090612 (PCT Application)
  CA 2727563 A 20101210 (PCT national entry)
Priority: IT 2008MI1079 A 20080613
Related Publication: WO 2009150533 A (Based on OPI patent )
Original IPC: G01N-33/15(B,I,H,EP,20060101,20110129,A,F)
Current IPC: G01N-33/15(B,I,H,EP,20060101,20110129,A,F)
EPO
Publication No. EP 2300817 A2 (Update 201124 E)
Publication Date: 20110330
**ANALYSE VON VI-SACCHARIDEN
 ANALYSIS OF VI SACCHARIDES
  ANALYSE DE SACCHARIDES VI**
Assignee: Novartis AG, Lichtstrasse 35, 4056 Basel, CH (NOVS)
Inventor: BERTI, Francesco, Novartis Vaccines, Via Fiorentina 1, I-53100
   Siena, IT
  MICOLI, Francesca, Novartis Vaccines, Via Fiorentina 1, I-53100 Siena, IT
  PROIETTI, Daniela, Novartis Vaccines, Via Fiorentina 1, I-53100 Siena, IT
Agent: Marshall, Cameron John, Carpmaels Ransford, One Southampton Row,
   London, WC1B 5HA, GB
Language: EN
Application: EP 2009762061 A 20090612 (Local application)
 WO 2009IB6087 A 20090612 (PCT Application)
Priority: IT 2008MI1079 A 20080613
Related Publication: WO 2009150533 A (Based on OPI patent )
Designated States: (Regional Confirmed) AT BE BG CH CY CZ DE DK EE ES FI FR
    GB GR HR HU IE IS IT LI LT LU LV MC MK MT NL NO PL PT RO SE SI SK TR AL
Original IPC: G01N-33/15(B,I,H,EP,20060101,20100107,A,F)
Current IPC: G01N-33/15(B,I,H,EP,20060101,20100107,A,F)
Original Abstract: ~Salmonella typhi Vi~ saccharide can be assayed in two
    new ways. First, its proton NMR spectrum can be used, with comparison
    to an internal Standard permitting quantitative analysis. Second, anion
    exchange chromatography with amperometric detection can be used on
   hydrolysed saccharide.
India
Publication No. IN 201004854 P2 (Update 201123 E)
Publication Date: 20110311
**Analysis of saccharides**
Assignee: NOVARTIS AG; CH (NOVS)
Inventor: BERTI F
 MICOLI F
 PROIETTI D
Language: EN
Application: IN 2010KN4854 A 20101220 (Local application)
  WO 2009IB6087 A 20090612 (PCT Application)
Priority: IT 2008MI1079 A 20080613
Original IPC: G01N-33/15(A)
Current IPC: G01N-33/15(A)
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WIPO
Publication No. WO 2009150533 A2 (Update 201008 B)
Publication Date: 20091217
**ANALYSIS OF VI SACCHARIDES
  ANALYSE DE SACCHARIDES VI**
Assignee: ~(except US)~ NOVARTIS AG, Lichtstrasse 35, CH-4056 Basel, CH
    Residence: CH Nationality: CH (NOVS)
  ~(only US)~ BERTI, Francesco, Novartis Vaccines, Via Fiorentina 1,
    I-53100 Siena, IT Residence: IT Nationality: IT
  ~(only US)~ MICOLI, Francesca, Novartis Vaccines, Via Fiorentina 1.
    I-53100 Siena, IT Residence: IT Nationality: IT
  ~(only US)~ PROIETTI, Daniela, Novartis Vaccines, Via Fiorentina 1,
    I-53100 Siena, IT Residence: IT Nationality: IT
Inventor: BERTI, Francesco, Novartis Vaccines, Via Fiorentina 1, I-53100
    Siena, IT Residence: IT Nationality: IT
  MICOLI, Francesca, Novartis Vaccines, Via Fiorentina 1, I-53100 Siena, IT
    Residence: IT Nationality: IT
  PROIETTI, Daniela, Novartis Vaccines, Via Fiorentina 1, I-53100 Siena, IT
    Residence: IT Nationality: IT
Agent: MARSHALL, Cameron, John et al., Carpmaels Ransford, 43-45
    Bloomsbury Square, London WC1A 2RA, GB
Language: EN (23 pages, 13 drawings)
Application: WO 2009IB6087 A 20090612 (Local application)
Priority: IT 2008MI1079 A 20080613
Designated States: (National Confirmed) AE AG AL AM AO AT AU AZ BA BB BG BH
    BR BW BY BZ CA CH CL CN CO CR CU CZ DE DK DM DO DZ EC EE EG ES FI GB GD
    GE GH GM GT HN HR HU ID IL IN IS JP KE KG KM KN KP KR KZ LA LC LK LR LS
    LT LU LY MA MD ME MG MK MN MW MX MY MZ NA NG NI NO NZ OM PE PG PH PL PT
    RO RS RU SC SD SE SG SK SL SM ST SV SY TJ TM TN TR TT TZ UA UG US UZ VC
    VN ZA ZM ZW
  (Regional Confirmed) AT BE BG CH CY CZ DE DK EE ES FI FR GB GR HR HU IE
    IS IT LT LU LV MC MK MT NL NO PL PT RO SE SI SK TR OA BW GH GM KE LS MW
    MZ NA SD SL SZ TZ UG ZM ZW EA
Original IPC: G01N-33/15(B,I,H,EP,20060101,A,F)
   G01N-33/15(B, I, M, 98, 20060101, C)
Current IPC: G01N-33/15(B,I,H,EP,20060101,20091217,A,F)
Current ECLA class: G01R-33/46
Current ECLA ICO class: S01N-224:352 S01N-333:255 S01N-400:10 S01R-330:400A
    S01R-330:404B
                   ~Salmonella typhi Vi~ saccharide can be assayed in two
Original Abstract:
    new ways. First, its proton NMR spectrum can be used, with comparison
    to an internal Standard permitting quantitative analysis. Second, anion
    exchange chromatography with amperometric detection can be used on
   hydrolysed saccharide.
   Le saccharide VI de ~Salmonella typhi~ peut etre dose de deux nouvelles
    facons. Tout d'abord, son spectre de RMN du proton peut etre utilise,
    avec comparaison avec un etalon interne permettant une analyse
    quantitative. Ensuite, une chromatographie d'echange d'anions avec une
    detection amperometrique peut etre utilisee sur un saccharide
    hydrolyse.
Publication No. WO 2009150533 A3 (Update 201010 E)
Publication Date: 20100204
**ANALYSIS OF VI SACCHARIDES**
Assignee: NOVARTIS AG; CH (NOVS)
Inventor: BERTI F, IT
 MICOLI F, IT
  PROIETTI D, IT
Language: EN
Application: WO 2009IB6087 A 20090612 (Local application)
Priority: IT 2008MI1079 A 20080613
Designated States: (National Confirmed) AE AG AL AM AO AT AU AZ BA BB BG BH
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BR BW BY BZ CA CH CL CN CO CR CU CZ DE DK DM DO DZ EC EE BG ES FI GB GD GE GH GM GT HN HR HU ID IL IN IS JP KE KG KM KN KP KR KZ LA LC LK LR LS LT LU LY MA MD ME MG MK MN MW MX MY MZ NA NG NI NO NZ OM PE PG PH PL PT RO RS RU SC SD SE SG SK SL SM ST SV SY TJ TM TN TR TT TZ UA UG US UZ VC VN ZA ZX ZW

(Regional Confirmed) AT BE BG CH CY CZ DE DK EE ES FI FR GB GR HR HU IE IS IT LT LU LV MC MK MT NL NO PL PT RO SE SI SK TR OA BW GH GM KE LS MW MZ NA SD SL SZ TZ UG ZM ZW EA

Original IPC: G01N-33/15(B,I,H,EP,20060101,A,F)

G01N-33/15(B,I,M,98,20060101,C)

Current IPC: G01N-33/15(B,I,H,EP,20060101,20100204,A,F)

Current ECLA class: G01R-33/46

Current ECLA ICO class: S01N-224:352 S01N-333:255 S01N-400:10 S01R-330:400A S01R-330:404B

Original Abstract: Salmonella typhi Vi saccharide can be assayed in two new ways. First, its proton NMR spectrum can be used, with comparison to an internal Standard permitting quantitative analysis. Second, anion exchange chromatography with amperometric detection can be used on hydrolysed saccharide.

10/7/5 (Item 5 from file: 351)

DIALOG(R)File 351:Derwent WPI

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0019309140

WPI ACC NO: 2009-L52824/200950

Membrane, useful for adsorption and removal of ****lipopolysaccharide**** from a suspension, comprises a polymeric substrate that binds to

****lipopolysaccharide****, which is from e.g. proteobacteria,

cyanobacteria and green sulfur bacteria
Patent Assignee: NOVARTIS AG (NOVS); BARBANI N (BARB-I); CIARDELLI G

(CIAR-I); COSTANTINO P (COST-I)
Inventor: BARBANI N; CIARDELLI G; COSTANTINO P; BRABANT N, CH; COSTANTINO

P, CH; SIADELY G, CH Patent Family (7 patents, 123 countries)

Patent Application Number Kind Date Number Kind Date Update WO 2009087571 A2 20090716 WO 2009IB133 A 20090107 200950 WO 2009087571 A3 20090903 WO 2009IB133 A 20090107 200958 E A 20090107 201072 EP 2244828 A2 20101103 EP 2009700597 A 20090107 WO 2009IB133 US 20100282684 A1 20101111 WO 2009IB133 A 20090107 201074 A 20100716 US 2010744306 CN 101909742 Α 20101208 CN 200980102069 A 20090107 201104 E WO 2009IB133 A 20090107 A1 20090716 CA 2711584 A 20090107 201110 E CA 2711584 WO 2009IB133 A 20090107 A 20100706 CA 2711584 JP 2011508772 TaT 20110317 WO 2009IB133 A 20090107 201121 E JP 2010541129 A 20090107

Priority Applications (no., kind, date): GB 2008228 A 20080107

Patent Details

Number Kind Lan Pg Dwg Filing Notes

WO 2009087571 A2 EN 13 2

National Designated States, Confirmed: AE AG AL AM AO AT AU AZ BA BB BG BH BR BM BY BZ CA CH CN CO CR CU CZ DE DK DM DO DZ EC EE EG ES FI GB GD GE GH GM GT HN HR HU ID IL IN 1S JP KE KG KM KN KP KR KZ LA LC LK LR LS LT LU LY MA MD ME MG MK MN MM MX MY MZ NA NG NI NO NZ OM PG PH PL PT RO RS RU SC SD SE SG SK SL SM ST SV SY TJ TM TN TR TT TZ UN UG US UZ VC VN ZA ZM ZW

Regional Designated States, Confirmed: AT BE BG CH CY CZ DE DK EE ES FI FR GB GR HR HU IE IS II LT LU LV MC MK MT NL NO PL PT RO SE SI SK TR OA BW GH GM KE LS MW MZ NA SD SL SZ TZ UG ZM ZW EA

WO 2009087571 A3 EN

National Designated States, Confirmed: AE AG AL AM AO AT AU AZ BA BB BG BH BR BW BY BZ CA CH CN CO CR CU CZ DE DK DM DO DZ EC EE EG ES FI GB GD GE GH GM GT HN HR HU ID IL IN 1S JP KE KG KM KN KP KR KZ LA LC LK LR LS LL LU LY MA MD ME MG MK MN MW MX MY MZ NA NG NI NO NZ OM PG PH PL PT RO RS RU SC SD SE SG SK SL SM ST SV SY TJ TM TN TR TT TZ UA UG US UZ VC VN ZA ZM ZW

Regional Designated States, Confirmed: AT BE BG CH CY CZ DE DK EE ES FI FR GB GR HR HU IE IS IT LIT UN LV MC MK MT NL NO PL PT RO SE SI SK TR OA BW GH GM KE LS MM MZ NA SD SL SZ TZ UG ZM ZW EA

EP 2244828 A2 EN PCT Application WO 2009IB133

Based on OPI patent WO 200908751 Regional Designated States, Confirmed: AT BE BG CH CY CZ DE DK EE ES FI FR BB GR HR HU IE IS IT LI LT LU LV MC MK MT NL NO PL PT RO SE SI SK TR AL BA RS

US 20100282684 A1 EN PCT Application WO 2009IB133 PCT Application WO 2009IB133 CN 101909742 A ZH Based on OPI patent WO 2009087571 CA 2711584 A1 EN PCT Application WO 2009IB133 PCT national entry CA 2711584 Based on OPI patent WO 2009087571 JP 2011508772 PCT Application WO 2009IB133 JΑ 1.5 Based on OPI patent WO 2009087571

Alerting Abstract WO A2

NOVELTY - Membrane comprises a polymeric substrate that binds ****lipopolysaccharide****.

DESCRIPTION - INDEPENDENT CLAIMS are included for:

- 1.a process for forming a polymeric substrate that binds to ****lipopolysaccharide****, comprising either contacting a homogeneous polymer solution and a template solution, carrying out a phase inversion of the resulting solution and removing the template, or contacting a monomer solution and a template solution, reacting crosslinking groups of the monomers to form a polymer and removing the template;
- 2.a method for the removal of ****lipopolysaccharide**** from a suspension comprising providing the polymeric substrate and contacting the suspension with the polymeric substrate; and
- 3.a polymeric substrate produced by the process.

USB - The membrane is useful for: adsorption of
****lipopolysaccharide****, where the ****lipopolysaccharide**** is from
Gram-negative bacteria (which are proteobacteria, cyanobacteria,
spirochetes, green sulfur bacteria, green non-sulfur bacteria,
crenarchaeota, cocci, bacilli or nosocomial bacteria); and removing
****lipopolysaccharide**** from a suspension (all claimed).

ADVANTAGE - The membrane selectively removes ****lipopolysaccharide****, or endotoxin, during the purification of molecules of biopharmaceutical interest.

Technology Focus

PHARMACEUTICALS - Preferred Components: The suspension comprises water and a pharmaceutical ingredient. The pharmaceutical ingredient is a bacterial vaccine.

POLYMERS - Preferred Components: The polymeric substrate is selective for at least one of heptose and 2-keto-3-deoxyoctonic acid. Preferred

Components: The template solution comprises at least one of heptose and Z-keto-3-deoxyoctonic acid. The polymeric substrate is in the form of a membrane or discrete particles. The polymeric substrate comprises one or more polar groups, preferably one or more hydroxyl groups. The polymeric substrate comprises poly(ethylene-co-vinyl alcohol). The ratio of ethylene:co-vinyl alcohol in the poly(ethylene-co-vinyl alcohol) is 30-60:70-40. Preferred Process: The process further comprises the step of making a membrane. The polymeric substrate is attached to a solid state support.

Title Terms/Index Terms/Additional Words: MEMBRANE; USEFUL; ADSORB; REMOVE; SUSPENSION; COMPRISE; POLYMERISE; SUBSTRATE; BIND; CYANOBACTERIA; GREEN; SULPHUR; BACTERIA

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Class Codes
International Classification (+ Attributes)
IPC + Level Value Position Status Version
  B01D-0015/04 A I F B 20060101
  B01D-0061/00 A I L B 20060101
  B01D-0067/00 A I L B 20060101
  B01J-0020/26 A I F B 20060101
  B01J-0020/28 A I L B 20060101
  C08F-0118/02 A I L B 20060101
  A61K-0039/00 A I L B 20060101
  A61K-0039/02 A I L B 20060101
A61K-0039/07 A I L B 20060101
A61K-0047/32 A I L B 20060101
A61K-0047/32 A I L B 20060101
B01D-0069/00 A I L B 20060101
  B01D-0071/38 A I L B 20060101
  B01D-0015/04 C I
                          B 20060101
  B01D-0061/00 C I L B 20090101
B01D-0061/00 C I B 20060101
  B01D-0067/00 C I L B 20090101
B01D-0067/00 C I B 20060101
  B01J-0020/22 C I F B 20090101
  B01J-0020/22 C I B 20060101
  B01J-0020/28 C I L B 20090101
  B01J-0020/28 C I
                         B 20060101
  C08F-0118/00 C I
                         B 20060101
ECLA: B01D-061/00, B01D-067/00K18D, B01J-020/26, B01J-020/28D24
ICO: L01D-323:24
US Classification, Current Main: 210-691000; Secondary: 210-690000,
526-319000
US Classification, Issued: 210691, 210690, 526319
JP Classification
  FI Term
                    Facet Rank Type
A61K-039/00
                    D
                            B secondary
A61K-039/02
                             A main
A61K-039/07
                             B secondary
A61K-047/32
                            B secondary
A61K-009/10
                             B secondary
B01D-069/00
                             B secondary
B01D-071/38
                             B secondary
F-Term View Point Additional
 Theme
        + Figure Code
 4C076
 4C085
 4D006
 40085
          AA03
 4C076
          AA22
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4C085
         BA07
 40085
          BA15
 40085
          CC07
 4C076
          CC31
 4C085
          DD37
 4C085
          EE01
 4C076
          EE03
 4C076
          EE06
 4C076
          FF70
 4D006
          GA01
 4C076
          GG43
 4D006
          MC34
 4D006
          NA54
 4D006
          PA01
 4D006
          PB02
 40006
          PB70
 4D006
          PC43
File Segment: CPI
DWPI Class: A18; A97; B04; J01; J04
Manual Codes (CPI/A-M): A03-A01; A10-E09B; A12-W11A; A12-W11L; B04-C02V;
 B04-C03; B14-S11B1; J01-C03; J04-X
Original Publication Data by Authority
Canada
Publication No. CA 2711584 Al (Update 201110 E)
Publication Date: 20090716
Assignee: NOVARTIS AG; CH (NOVS)
Inventor: COSTANTINO P, IT
 CIARDELLI G, IT
 BARBANI N. IT
Language: EN
Application: CA 2711584 A 20090107 (Local application)
  WO 2009IB133 A 20090107 (PCT Application)
 CA 2711584 A 20100706 (PCT national entry)
Priority: GB 2008228 A 20080107
Related Publication: WO 2009087571 A (Based on OPI patent )
Original IPC: B01D-61/00(B,I,H,EP,20060101,20100907,A,L)
    B01D-67/00(B,I,H,EP,20060101,20100907,A,L)
    B01J-20/26(B, I, H, EP, 20060101, 20100907, A, F)
   B01J-20/28(B, I, H, EP, 20060101, 20100907, A, L)
Current IPC: B01D-61/00(B,I,H,EP,20060101,20100907,A,L)
    B01D-67/00(B,I,H,EP,20060101,20100907,A,L)
    B01J-20/26(B,I,H,EP,20060101,20100907,A,F)
   B01J-20/28(B, I, H, EP, 20060101, 20100907, A, L)
China
Publication No. CN 101909742 A (Update 201104 E)
Publication Date: 20101208
**Lipopolysaccharide decontamination**
Assignee: NOVARTIS AG: CH (NOVS)
Inventor: COSTANTINO P. CH
 COSTANTINO PAOLO, CH
 SIADELY G, CH
 Siadely G, CH
 BRABANT N. CH
 BRABANT N, CH
Language: ZH
Application: CN 200980102069 A 20090107 (Local application)
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- WO 2009IB133 A 20090107 (PCT Application) Priority: GB 2008228 A 20080107
- Related Publication: WO 2009087571 A (Based on OPI patent)
- Original IPC: B01D-61/00(I,CN,20060101,A,L) B01D-67/00(I,CN,20060101,A,L) B01D-20/26(I,CN,20060101,A,F) B01D-20/28(I,CN,20060101,A,L)
- Current IPC: B01D-61/00(B,I,H,CN,20060101,20101209,A,L)
 - B01D-67/00 (B, I, H, CN, 20060101, 20101209, A, L)
 - B01J-20/26(B, I, H, CN, 20060101, 20101209, A, F)
- B01J-20/28(B, I, H, CN, 20060101, 20101209, A, L)
- Original Abstract: The invention claims materials and methods for the selective removal of lippoplysaccharide during the purification of molecules of biopharmaceutical interest, which are based on a polymeric substrate that binds lippoplysaccharide. Preferably, the polymeric substrate is selective for at least one of heptose and 2-keto-3-deoxyoctonic acid. The substrate can be formed by a process comprising: (i) contacting a homogeneous polymer solution and a template solution; (ii) carrying out a phase inversion of the resulting solution; and (iii) removing the template.
- Claim: [CLAIM 1] A membrane for adsorption of lipopolysaccharide, comprising a polymeric substrate that binds lipopolysaccharide.
 - [CLAIM 2] The membrane according to claim 1, wherein the polymeric substrate is selective for at least one of heptose and 2-keto-3-deoxyoctonic acid.
 - [CLAIM 3] The membrane according to either of claims 1 and 2, wherein the lipopolysaccharide is from Gram-negative bacteria.
 - [CLAIM 4] The membrane according to claim 3, wherein the Gram-negative bacteria are proteobacteria, cyanobacteria, spirochaetes, green sulphur bacteria, green non-sulphur bacteria, crenarchaeota, cocci, bacilli or nosocomial bacteria.
 - [CLAIM 5] A process for forming a polymeric substrate that binds lipopolysaccharide, comprising steps of: i. contacting a homogeneous polymer solution and a template solution; ii. carrying out a phase inversion of the resulting solution; and iii. removing the template.
 - [CLAIM 6] A process for forming a polymeric substrate that binds lipopolysaccharide, comprising steps of: i. contacting a monomer solution and a template solution; ii. reacting cross-linking groups of the monomers to form a polymer; and iii. removing the template.
 - [CLAIM 7] The process according to either of claims 5 and 6, further comprising the step of making a membrane.
 - [CLAIM 8] The process according to any of claims 5 to 7, wherein the template solution comprises at least one of heptose and 2-keto-3-deoxyoctonic acid.
 - [CLAIM 9] A method for the removal of lipopolysaccharide from a suspension comprising steps of: i. providing a polymeric substrate that binds lipopolysaccharide; and ii. contacting the suspension with the polymeric substrate.
 - [CLAIM 10] The method according to claim 9, wherein the polymeric substrate is in the form of a membrane.
 - [CLAIM 11] The method according to claim 9, wherein the polymeric substrate is in the form of discrete particles.
 - [CLAIM 12] The method according to claim 9, wherein the polymeric
 - substrate is attached to a solid state support. [CLAIM 13] The method according to any one of claims 9 to 12, wherein the polymeric substrate is selective for at least one of heptose and 2-keto-3-decoxvoctonic acid.
 - [CLAIM 14] The method according to any one of claims 9 to 13, wherein the lipopolysaccharide is from Gram-negative bacteria.
 - [CLAÎN 15] The method according to claim 14, wherein the Gram-negative bacteria are protoobacteria, cyanobacteria, spirochaetes, green sulphur bacteria, green non-sulphur bacteria, crenarchaeota or nosocomial bacteria.
 - [CLAIM 16] The method according to any one of claims 9 to 15, wherein the

suspension comprises water. [CLAIM 17] The method according to any one of claims 9 to 16, wherein the suspension comprises a pharmaceutical ingredient. [CLAIM 18] The method according to claim 17, wherein the pharmaceutical ingredient is a bacterial vaccine. [CLAIM 19] The membrane, process or method according to any preceding claim, wherein the polymeric substrate comprises one or more polar groups. [CLAIM 20] The membrane, process or method according to claim 19, wherein the polymeric substrate comprises one or more hydroxyl groups. [CLAIM 21] The membrane, process or method according to any preceding claim, wherein the polymeric substrate comprises poly(ethylene-co-vinyl alcohol). [CLAIM 22] The membrane, process or method according to claim 21, wherein the ratio of ethylene: co-vinyl alcohol in the poly(ethylene-co-vinyl alcohol) is 30-60:70-40. [CLAIM 23] A polymeric substrate produced by the process according to any one of claims 5 to 7. Publication No. EP 2244828 A2 (Update 201072 E) Publication Date: 20101103 **LIPOPOLYSACCHARID-DEKONTAMINATION LIPOPOLYSACCHARIDE DECONTAMINATION DECONTAMINATION DE LIPOPOLYSACCHARIDE** Assignee: Novartis AG, Lichtstrasse 35, 4056 Basel, CH (NOVS) Inventor: COSTANTINO, Paolo, Novartis VaccinesDiagnostics, Via Fiorentina 1, I-53100 Siena, IT Agent: Marshall, Cameron John, Carpmaels Ransford, One Southampton Row, London, WC1B 5HA, GB Language: EN Application: EP 2009700597 A 20090107 (Local application) WO 2009IB133 A 20090107 (PCT Application) Priority: GB 2008228 A 20080107 Related Publication: WO 2009087571 A (Based on OPI patent) Designated States: (Regional Confirmed) AT BE BG CH CY CZ DE DK EE ES FI FR GB GR HR HU IE IS IT LI LT LU LV MC MK MT NL NO PL PT RO SE SI SK TR AL BA RS Original IPC: B01D-61/00(B,I,H,EP,20060101,20090804,A,L) B01D-61/00(B,I,M,98,20060101,20090804,C) B01D-67/00(B,I,H,EP,20060101,20090804,A,L) B01D-67/00(B,I,M,98,20060101,20090804,C) B01J-20/22(B, I, M, 98, 20060101, 20090804, C) B01J-20/26(B, I, H, EP, 20060101, 20090804, A, F) B01J-20/28(B, I, H, EP, 20060101, 20090804, A, L) B01J-20/28(B, I, M, 98, 20060101, 20090804, C) Current IPC: B01D-61/00(B,I,H,EP,20060101,20090804,A,L) B01D-61/00(B, I, M, 98, 20060101, 20090804, C) B01D-67/00(B,I,H,EP,20060101,20090804,A,L) B01D-67/00(B, I, M, 98, 20060101, 20090804, C) B01J-20/22(B, I, M, 98, 20060101, 20090804, C) B01J-20/26(B, I, H, EP, 20060101, 20090804, A, F) B01J-20/28(B, I, H, EP, 20060101, 20090804, A, L) B01J-20/28(B, I, M, 98, 20060101, 20090804, C) Original Abstract: Materials and methods for the selective removal of lipopolysaccharide during the purification of molecules of biopharmaceutical interest are based on a polymeric substrate that binds lipopolysaccharide. Preferably, the polymeric substrate is selective for at least one of heptose and 2-keto-3-deoxyoctonic acid. The substrate can be formed by a process comprising: (i) contacting a homogeneous polymer solution and a template solution; (ii) carrying out a phase inversion of the resulting solution; and (iii) removing the

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template.
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Japan
Publication No. JP 2011508772 W (Update 201121 E)
Publication Date: 20110317
Language: JA (15 pages)
Application: JP 2010541129 A 20090107 (Local application)
  WO 2009IB133 A 20090107 (PCT Application)
Priority: GB 2008228 A 20080107
Related Publication: WO 2009087571 A (Based on OPI patent )
Original IPC: A61K-39/00(B,I,H,JP,20060101,20110218,A,L)
    A61K-39/02(B, I, H, JP, 20060101, 20110218, A, F)
    A61K-39/07(B, I, H, JP, 20060101, 20110218, A, L)
    A61K-47/32(B, I, H, JP, 20060101, 20110218, A, L)
    A61K-9/10(B, I, H, JP, 20060101, 20110218, A, L)
    B01D-69/00(B, I, H, JP, 20060101, 20110218, A, L)
    B01D-71/38(B, I, H, JP, 20060101, 20110218, A, L)
Current IPC: A61K-39/00(B,I,H,JP,20060101,20110218,A,L)
    A61K-39/02(B, I, H, JP, 20060101, 20110218, A, F)
    A61K-39/07(B, I, H, JP, 20060101, 20110218, A, L)
    A61K-47/32(B, I, H, JP, 20060101, 20110218, A, L)
    A61K-9/10(B, I, H, JP, 20060101, 20110218, A, L)
    B01D-69/00(B, I, H, JP, 20060101, 20110218, A, L)
    B01D-71/38(B, I, H, JP, 20060101, 20110218, A, L)
Current JP FI-Terms: A61K-39/02 (main, A) A61K-39/00 D (secondary, B)
    A61K-39/07 (secondary, B) A61K-47/32 (secondary, B) A61K-9/10
    (secondary, B) B01D-69/00 (secondary, B) B01D-71/38 (secondary, B)
Current JP F-Terms: 4C076 4C085 4D006 4C085AA03 4C076AA22 4C085BA07
    4C085BA15 4C085CC07 4C076CC31 4C085DD37 4C085EE01 4C076EE03 4C076EE06
    4C076FF70 4D006GA01 4C076GG43 4D006MC34 4D006NA54 4D006PA01 4D006PB02
    4D006PB70 4D006PC43
United States
Publication No. US 20100282684 Al (Update 201074 E)
Publication Date: 20101111
**LIPOPOLYSACCHARIDE DECONTAMINATION**
Assignee: Costantino, Paolo, Siena, IT Residence: IT (COST-I)
  Ciardelli, Gianluca, Pisa, IT Residence: IT (CIAR-I)
  Barbani, Niccoletta, Pisa, IT Residence: IT (BARB-I)
Inventor: Costantino, Paolo, Siena, IT Residence: IT
  Ciardelli, Gianluca, Pisa, IT Residence: IT
 Barbani, Niccoletta, Pisa, IT Residence: IT
Agent: NOVARTIS VACCINES AND DIAGNOSTICS INC., INTELLECTUAL PROPERTY-
    X100B, P.O. BOX 8097, Emeryville, CA, US
Language: EN
Application: US 2010744306 A 20100716 (Local application)
  WO 2009IB133 A 20090107 (PCT Application)
Priority: GB 2008228 A 20080107
Original IPC: B01D-15/04(B,I,H,US,20060101,20101111,A,F)
    B01D-15/04(B, I, M, 98, 20060101, 20101111, C)
    C08F-118/00(B, I, M, 98, 20060101, 20101111, C)
    C08F-118/02(B, I, H, US, 20060101, 20101111, A, L)
Current IPC: B01D-15/04(B,I,H,US,20060101,20101111,A,F)
    B01D-15/04(B, I, M, 98, 20060101, 20101111, C)
    C08F-118/00(B, I, M, 98, 20060101, 20101111, C)
    C08F-118/02(B, I, H, US, 20060101, 20101111, A, L)
Current ECLA class: B01D-61/00 B01D-67/00K18D B01J-20/26 B01J-20/28D24
Current ECLA ICO class: L01D-323:24
Current US Class (main): 210-691000
Current US Class (secondary): 210-690000 526-319000
Original US Class (main): 210691
Original US Class (secondary): 210690 526319
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Original Abstract: Materials and methods for the selective removal of
    lipopolysaccharide during the purification of molecules of
    bio-pharmaceutical interest are based on a polymeric substrate that
   binds lipopolysaccharide. Preferably, the polymeric substrate is
    selective for at least one of heptose and 2-keto-3-deoxyoctonic acid.
    The substrate can be formed by a process comprising: (i) contacting a
    homogeneous polymer solution and a template solution; (ii) carrying out
    a phase inversion of the resulting solution; and (iii) removing the
    template.
Claim:
**1**. A membrane for adsorption of lipopolysaccharide, comprising a
        polymeric substrate that binds lipopolysaccharide.
WIPO
Publication No. WO 2009087571 A2 (Update 200950 B)
Publication Date: 20090716
**LIPOPOLYSACCHARIDE DECONTAMINATION
  DECONTAMINATION DE LIPOPOLYSACCHARIDE**
Assignee: ~(except US)~ NOVARTIS AG, Lichtstrasse 35, CH-4056 Basle, CH
    Residence: CH Nationality: CH (NOVS)
  ~(only US)~ COSTANTINO, Paolo, Novartis Vaccines Diagnostics, Via
    Fiorentina 1, I-53100 Siena, IT Residence: IT Nationality: IT
Inventor: COSTANTINO, Paolo, Novartis Vaccines Diagnostics, Via Fiorentina
    1, I-53100 Siena, IT Residence: IT Nationality: IT
Agent: MARSHALL, Cameron, John, Carpmaels Ransford, 43-45 Bloomsbury
    Square, London WC1A 2RA, GB
Language: EN (13 pages, 2 drawings)
Application: WO 2009IB133 A 20090107 (Local application)
Priority: GB 2008228 A 20080107
Designated States: (National Confirmed) AE AG AL AM AO AT AU AZ BA BB BG BH
    BR BW BY BZ CA CH CN CO CR CU CZ DE DK DM DO DZ EC EE EG ES FI GB GD GE
    GH GM GT HN HR HU ID IL IN IS JP KE KG KM KN KP KR KZ LA LC LK LR LS LT
    LU LY MA MD ME MG MK MN MW MX MY MZ NA NG NI NO NZ OM PG PH PL PT RO RS
    RU SC SD SE SG SK SL SM ST SV SY TJ TM TN TR TT TZ UA UG US UZ VC VN ZA
    ZM ZW
  (Regional Confirmed) AT BE BG CH CY CZ DE DK EE ES FI FR GB GR HR HU IE
    IS IT LT LU LV MC MK MT NL NO PL PT RO SE SI SK TR OA BW GH GM KE LS MW
    MZ NA SD SL SZ TZ UG ZM ZW EA
Original IPC: B01D-61/00(B,I,H,EP,20060101,A,L)
    B01D-61/00(B,I,M,98,20060101,C) B01D-67/00(B,I,H,EP,20060101,A,L)
    B01D-67/00(B,I,M,98,20060101,C) B01J-20/22(B,I,M,98,20060101,C)
    B01J-20/26(B,I,H,EP,20060101,A,F) B01J-20/28(B,I,H,EP,20060101,A,L)
    B01J-20/28(B, I, M, 98, 20060101, C)
Current IPC: B01D-61/00(B,I,H,EP,20060101,20090716,A,L)
    B01D-61/00(B,I,H,EP,20090101,20090716,C,L)
    B01D-67/00(B,I,H,EP,20060101,20090716,A,L)
    B01D-67/00(B,I,H,EP,20090101,20090716,C,L)
    B01J-20/22(B,I,H,EP,20090101,20090716,C,F)
    B01J-20/26(B, I, H, EP, 20060101, 20090716, A, F)
    B01J-20/28(B, I, H, EP, 20060101, 20090716, A, L)
    B01J-20/28(B, I, H, EP, 20090101, 20090716, C, L)
Current ECLA class: B01D-61/00 B01D-67/00K18D B01J-20/26 B01J-20/28D24
Current ECLA ICO class: L01D-323:24
Original Abstract: Materials and methods for the selective removal of
    lipopolysaccharide during the purification of molecules of
    biopharmaceutical interest are based on a polymeric substrate that
    binds lipopolysaccharide. Preferably, the polymeric substrate is
   selective for at least one of heptose and 2-keto-3-deoxyoctonic acid.
    The substrate can be formed by a process comprising: (i) contacting a
    homogeneous polymer solution and a template solution; (ii) carrying out
    a phase inversion of the resulting solution; and (iii) removing the
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template.
   L'invention concerne des matieres et des procedes pour le retrait
    selectif de lipopolysaccharide durant la purification de molecules
    d'interet biopharmaceutique qui sont fondes sur un substrat polymere
    qui se lie au lipopolysaccharide. De preference, le substrat polymere
    est selectif pour au moins l'un parmi l'heptose et l'acide
    2-ceto-3-desoxyoctonique. Le substrat peut etre forme par un procede
    consistant a: (i) mettre en contact une solution polymere homogene et
    une solution modele; (ii) a realiser une inversion de phase de la
    solution resultante; et (iii) a retirer le modele.
Publication No. WO 2009087571 A3 (Update 200958 E)
Publication Date: 20090903
**LIPOPOLYSACCHARIDE DECONTAMINATION**
Assignee: NOVARTIS AG; CH (NOVS)
Inventor: BARBANI N. IT
 CIARDELLI G, IT
  COSTANTINO P, IT
Language: EN
Application: WO 2009IB133 A 20090107 (Local application)
Priority: GB 2008228 A 20080107
Designated States: (National Confirmed) AE AG AL AM AO AT AU AZ BA BB BG BH
    BR BW BY BZ CA CH CN CO CR CU CZ DE DK DM DO DZ EC EE EG ES FI GB GD GE
    GH GM GT HN HR HU ID IL IN IS JP KE KG KM KN KP KR KZ LA LC LK LR LS LT
    LU LY MA MD ME MG MK MN MW MX MY MZ NA NG NI NO NZ OM PG PH PL PT RO RS
    RU SC SD SE SG SK SL SM ST SV SY TJ TM TN TR TT TZ UA UG US UZ VC VN ZA
    ZM ZW
  (Regional Confirmed) AT BE BG CH CY CZ DE DK EE ES FI FR GB GR HR HU IE
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    MZ NA SD SL SZ TZ UG ZM ZW EA
Original IPC: B01D-61/00(B,I,H,EP,20060101,A,L)
    B01D-61/00(B,I,M,98,20060101,C) B01D-67/00(B,I,H,EP,20060101,A,L)
    B01D-67/00(B,I,M,98,20060101,C) B01J-20/22(B,I,M,98,20060101,C)
    B01J-20/26(B,I,H,EP,20060101,A,F) B01J-20/28(B,I,H,EP,20060101,A,L)
    B01J-20/28(B, I, M, 98, 20060101, C)
Current IPC: B01D-61/00(B,I,H,EP,20060101,20090903,A,L)
   B01D-61/00(B, I, H, EP, 20090101, 20090903, C, L)
    B01D-67/00(B, I, H, EP, 20060101, 20090903, A, L)
    B01D-67/00(B, I, H, EP, 20090101, 20090903, C, L)
    B01J-20/22(B,I,H,EP,20090101,20090903,C,F)
    B01J-20/26(B, I, H, EP, 20060101, 20090903, A, F)
    B01J-20/28(B, I, H, EP, 20060101, 20090903, A, L)
    B01J-20/28(B, I, H, EP, 20090101, 20090903, C, L)
Current ECLA class: B01D-61/00 B01D-67/00K18D B01J-20/26 B01J-20/28D24
Current ECLA ICO class: L01D-323:24
Original Abstract: Materials and methods for the selective removal of
    lipopolysaccharide during the purification of molecules of
    biopharmaceutical interest are based on a polymeric substrate that
    binds lipopolysaccharide. Preferably, the polymeric substrate is
    selective for at least one of heptose and 2-keto-3-deoxyoctonic acid.
    The substrate can be formed by a process comprising: (i) contacting a
    homogeneous polymer solution and a template solution; (ii) carrying out
    a phase inversion of the resulting solution; and (iii) removing the
    template.
 10/7/6
            (Item 6 from file: 351)
DIALOG(R)File 351:Derwent WPI
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0019248030

WPI ACC NO: 2009-L02218/200946

Cultivating Streptococcus for ****capsular**** ****polysaccharide**** (cps) production comprises providing an inoculum of a strain of Streptococcus expressing the cps, and cultivating the strain by fermentation

Patent Assignee: NOVARTIS AG (NOVS); NOVARTIS VACCINES&DIAGNOSTICS INC

Inventor: BAZZOCCHI G, BERTI F, CICALA C, CICALA C M, CONSTANTINO P; COSTANTINO P; FONTANI S; NORBELLI F; OLIVIERI R; BAZZOCCHI G, CH; BERTI F, CH; CICALA C M, CH; COSTANTINO P, CH; FONTANI S, CH; NORELLI F, CH; OLIVIERI R, CH

Patent Family (9 patents, 123 countries) Application Number Kind Date Number Kind Date Update WO 2009081276 A2 20090702 WO 2008IB3729 A 20081219 200946 WO 2009081276 A3 20090903 WO 2008IB3729 A 20081219 200958 E AU 2008339553 A1 20090702 AU 2008339553 A 20081219 201050 E CA 2708878 A1 20090702 CA 2708878 A 20081219 201064 E WO 2008IB3729 A 20081219 A 20100610 CA 2708878 EP 2235159 A2 20101006 EP 2008864247 A 20081219 201065 E WO 2008IB3729 A 20081219 20101028 US 20100272755 A1 US 20078941 P 20071220 201071 E WO 2008IB3729 A 20081219 US 2010747914 A 20100613 IN 201002275 P2 20101008 WO 2008IB3729 A 20081219 201072 IN 2010KN2275 A 20100622 CN 101932698 Α 20101229 CN 200880126144 A 20081219 201109 A WO 2008TB3729 20081219 WO 2008IB3729 JP 2011507501 20110310 A 20081219 201118 JP 2010538947 A 20081219

Priority Applications (no., kind, date): US 20078941 P 20071220; US 20078941 P 20071220; GB 200818453 A 20081008

Patent Details

Number Kind Lan

WO 2009081276 A2 EN 159 30

WO 2009081276 AZ EN 159 30

National Designated States, Confirmed: AE AG AL AM AO AT AU AZ BA BB BG BH BR BW BY BZ CA CH CN CO CR CU CZ DE DK DM DD DZ BC EE EG ES FIG BG BG GH GM GT HN HR HU ID IL IN IS JP KE KG KW KN KP KR KZ LA LC LK LR LS LT LU LY MA MD ME MG MK MN MW KM YM ZN AN GN IN NON ZO MP EO HP LP FT ROS RU SC SD SE SG SK SL SM ST SV SY TJ TM TN TR TT TZ UA UG US UZ VC VN ZA ZM ZW

Pa Dwa Filina Notes

Regional Designated States, Confirmed: AT BE BG CH CY CZ DE DK EE ES FI FR GB GR HR HU IE IS IT LT LU LV MC MT NL NO PL PT RO SE SI SK TR OA BW GH GM KE LS MW MZ NA SD SL SZ TZ UG ZM ZW EA

WO 2009081276 A3 EN

National Designated States, Confirmed: AE AG AL AM AO AT AU AZ BA BB BG BH BR W BY BZ CA CH CN CO CR CU CZ DE DK DM DO DZ EC EE EG ES FI GB GD GE GH GH GT HN HR HU ID IL IN IS JP KE KG KM KN KP KR KZ LA LC LK LR LS LL LU LY MA MD ME MG MK MN MM MX MY MZ NA NG NI NO NZ OM PG PH PL PI RO RS RU SC SD SE SG SK SL SM ST SV SY IJ TM TN TR TT TZ UA UG US UZ VC VN ZA ZM ZM

Regional Designated States, Confirmed: AT BE BG CH CY CZ DE DK EE ES FI FR
GB GR HR HU IE IS IT LT LU LV MC MT NL NO PL PT RO SE SI SK TR OA BW GH
GM KE LS MW MZ NA SD SL SZ TZ UG ZM ZW EA

AU 2008339553 Al EN Based on OPI patent WO 2009081276 CA 2708878 Al EN PCT Application WO 20081B3729

PCT national entry CA 2708878
Based on OPI patent WO 2009081276
EP 2235159 A2 EN PCT Application WO 20081B3729
Based on OPI patent WO 2009081276

Regional Designated States, Confirmed: AT BE BG CH CY CZ DE DK EE ES FI FR

GB GR HR HU IE IS IT LI LT LU LV MC MT NL NO PL PT RO SE SI SK TR AL BA

US 20100272755 Al EN Related to Provisional US 20078941 PCT Application WO 2008IB3729 IN 201002275 P2 EN PCT Application WO 2008IB3729 A ZH PCT Application WO 2008IB3729 CN 101932698 Based on OPI patent WO 2009081276 JP 2011507501 W JA 120 PCT Application WO 2008IB3729 Based on OPI patent WO 2009081276

Alerting Abstract WO A2

NOVELTY — Cultivating "Streptococcus" for production of cps on a mannfacturing scale comprises providing an inoculum of a strain of "Streptococcus" expressing the cps, and cultivating the strain by fermentation, where the cultivating comprises a linear addition of a carbon source to a cultivating medium, and does not use an algorithm to control the cultivating by monitoring a pH of the cultivating medium.

DESCRIPTION - INDEPENDENT CLAIMS are: (1) a cultivating medium comprising a ~Streptococcus ~ strain, a phosphate source, a carbon source, a vitamin source, and an amino acid source to grow ~Streptococcus ~ , where the vitamin source consists of six or fewer vitamins selected from the following list of seven vitamins: biotin, niacinamide, calcium pantothenate, riboflavin, thiamine hydrochloride, pyridoxine hydrochloride and folic acid, where two of the vitamins have to be calcium pantothenate and niacinamide; (2) a cultivating medium comprising a ~Streptococcus strain, a yeast extract, a phosphate source, a carbon source, a vitamin source, and optionally an amino acid source to grow ~Streptococcus ~ , where the vitamin source consists of four or fewer vitamins selected from the following list of five vitamins: biotin, niacinamide, riboflavin, thiamine hydrochloride and pyridoxine hydrochloride, where one of the vitamins has to be biotin; #a method for purifying a cps from ~Streptococcus agalactiae ~ by a step of filtration using an adherent filter; and (3) a method for producing a purified cps by providing a crude isolate containing a cps, removing an alcohol precipitate formed by contacting the crude isolate with an alcohol solution, filtering to remove smaller molecular weight compounds while retaining the cps, and removing protein contaminants with a protein adherent filter to produce the purified cps.

ACTIVITY - Virucide; Antiinflammatory; Respiratory-Gen; Immunostimulant. No biological data given.

MECHANISM OF ACTION - Vaccine.

USE - The method and medium are useful for cultivating "Streptococcus" for production of cps (claimed). The cps and composition are used in therapy, e.g. for manufacturing a medicament for the treatment of disease, preferably the disease is influenza or pneumonia, and for eliciting systemic and/or mucosal immunity.

ADVANTAGE - The present invention provides a simplified purification procedure that will produce higher levels of purity with fewer complicated and/or expensive purification steps. It provides a purification procedure that provides a good yield of cps whatever the initial purity of the ****polysaccharide****.

Technology Focus

BIOTECHNÓLOGY - Preferred Method: The method for cultivating -Streptococcus ~ for production of cps on a manufacturing scale further comprises recovering the cps. The strain of ~Streptococcus ~ further comprises S. avalactica s ~, where the strain of S. agalactica is 090, H36b, CBJIII, or M781. An optical density (OD) of the inoculum is 0.6-1.8. A pH of the cultivating medium is 6.0-7.5, preferably 7.3. A temperature of the cultivating medium is 34-38 (deg) C, preferably 36 (deg) C. The carbon source further comprises glucose. The cultivation further comprises monitoring an OD of the cultivating medium such that when the OD reaches a

designated level, the linear addition of a carbon source is initiated, where the designated level is selected to achieve a higher volumetric production of cps. The designated level is 9.8-10.2, preferably 10. The cultivation further comprises monitoring an OD of the cultivating medium such that when the OD reaches a first and second instantaneous addition level, two instantaneous additions of yeast extract are initiated prior to the linear addition of a carbon source. The first and second instantaneous addition levels are selected to achieve a higher volumetric production of cps. The first instantaneous addition level is 2.8-3.2, preferably 3.0. The second instantaneous addition level is 4.3-4.7, preferably 4.5. The cultivating medium is a defined medium comprising a phosphate source, a mineral source, a carbon source, a vitamin source, and an amino acid source to grow ~Streptococcus ~ , where the vitamin source consists of six or fewer vitamins selected from the following list of seven vitamins: biotin, niacinamide, calcium pantothenate, riboflavin, thiamine hydrochloride, pyridoxine hydrochloride and folic acid, where two of the vitamins have to be calcium pantothenate and niacinamide. The cultivating medium is a complex medium comprising a yeast extract, a phosphate source, a carbon source, a vitamin source, and optionally an amino acid source to grow ~Streptococcus ~ , where the vitamin source consists of four or fewer vitamins selected from the following list of five vitamins: biotin, niacinamide, riboflavin, thiamine hydrochloride and pyridoxine hydrochloride, where one of the vitamins has to be biotin. The method for purifying a cps from ~Streptococcus ~ agalactiae does not include a step of cationic detergent treatment to precipitate the cps followed by a step of re-solubilization of the cps, where the adherent filter is a protein adherent filter, and where the adherent filter is a carbon filter. The step of filtration using an adherent filter is preceded by (i) alcoholic precipitation of contaminating proteins and/or nucleic acids, and (ii) diafiltration. The step of filtration using an adherent filter is followed by (iv) re-N-acetylation, and (v) diafiltration. The method for producing purified cps further comprises re-N-acetylating the purified cps, precipitating the purified cps, and formulating a vaccine with the cps as a component. The removing step comprises addition of an alcohol solution to a concentration sufficient to precipitate nucleic acid contaminants but not the cps, where the alcohol solution comprises ethanol. The alcohol solution further comprises calcium chloride (CaCl 2). The alcohol solution is added to a concentration of 10-50% ethanol, preferably 30% ethanol. The protein adherent filter is an activated carbon filter. Preferred Medium: In the cultivating medium of (1), the phosphate source consists of potassium hydrogen phosphate (K 2 HPO 4), potassium dihydrogen phosphate (KH 2 PO 4), sodium hydrogen phosphate monohydrate (Na 2 HPO 4 .H 2 O), sodium dihydrogen phosphate monohydrate (NaH 2 PO 4 .H 2 O), or sodium chloride (NaCl). The carbon source is glucose. The vitamin source consists of 3-6, or fewer from the following list of seven vitamins: biotin, niacinamide, calcium pantothenate, riboflavin, thiamine hydrochloride, pyridoxine hydrochloride, and folic acid, where two have to be calcium pantothenate and niacinamide. The vitamin source consists of calcium pantothenate and niacinamide. The amino acid source consists of 16-19 or fewer from the following list of nineteen amino acids: alanine, arginine, glutamine, glycine, histidine, isoleucine, leucine, lysine, methionine, phenylalanine, proline, serine, threonine, tryptophan, valine, aspartic acid, cysteine hydrochloride, glutamic acid, and tyrosine, where fifteen have to be arginine, glycine, histidine, isoleucine, leucine, lysine, methionine, phenylalanine, serine, threonine, tryptophan, valine, cysteine hydrochloride, glutamic acid, and tyrosine. In the cultivating medium of (2), the vitamin source consists of 2-4 or fewer vitamins selected from the following list of five vitamins: biotin, niacinamide, riboflavin, thiamine hydrochloride and pyridoxine hydrochloride, where one of the vitamins has to be biotin. The vitamin source consists of biotin.

; ****POLYSACCHARIDE****; PRODUCE; COMPRISE; INOCULATE; STRAIN; EXPRESS; FERMENTATION

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Class Codes
International Classification (Main): C12N-001/20
 (Additional/Secondary): C12P-019/04
International Classification (+ Attributes)
IPC + Level Value Position Status Version
 A61K-0039/04 A I F B 20060101
 A61K-0039/04 A I L B 20060101
 A61K-0039/04 A I L
                           20060101
 A61K-0039/09 A I F B 20060101
  C07H-0001/00 A I L B 20060101
  C12N-0001/20 A I F B 20060101
  C12N-0001/20 A I L B 20060101
  C12N-0001/20 A I F
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  C12P-0019/04 A I L B 20060101
  C12P-0019/04 A I L
                           20060101
  C12Q-0003/00 A I L B 20060101
  A61K-0039/00 A I L B 20060101
  A61K-0039/39 A I L B 20060101
  A61P-0031/04 A I L B 20060101
  A61K-0039/04 C I L B 20090101
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A61K-0039/09 C I
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 C12N-0001/20 C I F B 20090101
  C12N-0001/20 C I L B 20090101
 C12N-0001/20 C I
                       B 20060101
 C12P-0019/00 C I L B 20090101
 C12P-0019/00 C I B 20060101
 C12Q-0003/00 C I
                      B 20060101
ECLA: C08B-037/00P, C12N-001/20, C12P-019/04
ICO: K61K-039:00
US Classification, Current Main: 424-244100; Secondary: 435-003000,
435-101000, 435-253400, 536-123100
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 FI Term
                  Facet Rank Type
A61K-039/00
                  G
                        B secondary
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A61P-031/04
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C12N-001/20
                 A
                         A main
C12N-001/20
                         C linked
                 A
C12P-019/04
                 С
                         B secondary
C12P-019/04
                 C
                         C linked
C12R-001/46
                         C linked
F-Term View Point Additional
 Theme
       + Figure Code
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BARR

BB02

4B065

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DWPI Class: B04; D16; T01
Manual Codes (EPI/S-X): T01-J06A: T01-J13A
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  B05-A01B; B05-B02A3; B05-C07; B06-D01; B06-D09; B06-F03; B07-D03;
  B07-D04C; B07-D09; B10-A07A; B10-A17; B10-B01; B10-B02; B14-A02B2;
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Language: EN
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Related Publication: WO 2009081276 A (Based on OPI patent )
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    C12N-1/20(B, I, H, EP, 20060101, 20090903, A, L)
    C12P-19/04(B, I, H, EP, 20060101, 20090903, A, L)
Current IPC: A61K-39/04(B,I,H,EP,20060101,20090903,A,F)
    A61K-39/04(B,I,H,EP,20090101,20090903,C,F)
    C12N-1/20(B, I, H, EP, 20060101, 20090903, A, L)
    C12N-1/20(B, I, H, EP, 20090101, 20090903, C, L)
    C12P-19/00(B, I, H, EP, 20090101, 20090903, C, L)
    C12P-19/04(B, I, H, EP, 20060101, 20090903, A, L)
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Canada
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Publication Date: 20090702
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Application: CA 2708878 A 20081219 (Local application)
  WO 2008IB3729 A 20081219 (PCT Application)
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Priority: US 20078941 P 20071220
  GB 200818453 A 20081008
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Original IPC: A61K-39/04(B,I,H,EP,20060101,20100809,A,L)
    A61K-39/04(B, I, M, 98, 20060101, 20100809, C)
    C12N-1/20(B, I, H, EP, 20060101, 20100809, A, F)
    C12N-1/20(B, I, M, 98, 20060101, 20100809, C)
    C12P-19/00(B, I, M, 98, 20060101, 20100809, C)
    C12P-19/04(B, I, H, EP, 20060101, 20100809, A, L)
Current IPC: A61K-39/04(B,I,H,EP,20060101,20100809,A,L)
    A61K-39/04(B, I, M, 98, 20060101, 20100809, C)
    C12N-1/20(B, I, H, EP, 20060101, 20100809, A, F)
    C12N-1/20(B, I, M, 98, 20060101, 20100809, C)
    C12P-19/00(B, I, M, 98, 20060101, 20100809, C)
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China
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**Fermentation processes for cultivating streptococci and purification
    processes for obtaining cps therefrom**
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Current IPC: A61K-39/04(I,CN,20060101,A,L) C12N-1/20(I,CN,20060101,A,F) C12P-19/04(I,CN,20060101,A,L)

Original Abstract: This invention is in the field of bacterial cultures, and specifically relates to the optimization of culture conditions to improve the production of bacterial capsular polysaccharides from Streptococcus strains in fed batch culture and to novel purification methods suitable for production scale purification of bacterial capsular polysaccharides from Streptococcus strains resulting in higher levels of purity than previously obtained for production scale.

Claim: [CLAIM 1] A method for cultivating Streptococcus for production of capsular polysaccharides (cps) on a manufacturing scale, wherein said method comprises (a) providing an inoculum of a strain of Streptococcus expressing the cps, and (b) cultivating the strain by fermentation, wherein said cultivating comprises a linear addition of a carbon source to a cultivating medium, and does not use an algorithm to control the cultivating by monitoring a pH of the cultivating medium.

[CLAIM 2] The method according to claim 1 further comprising step (c) recovering the capsular polysaccharide.

[CLAIM 3] The method according to claim 1 or claim 2, wherein said strain of Streptococcus further comprises Streptococcus agalactiae.

[CLAIM 4] The method according to claim 3, wherein said strain of Streptococcus agalactiae is 090, H36b, CBJ111, or M781.

[CLAIM 5] The method according to any one of claims 1-4, wherein an optical density (OD) of the inoculum is between about 0.6 and about 1.8.

[CLAIM 6] The method according to any one of claims 1-5, wherein a pH of the cultivating medium is between about 6.0 and about 7.5.

[CLAIM 7] The method according to claim 6, wherein the pH is about 7.3. [CLAIM 8] The method according to any one of claims 1-7, wherein a

temperature of the cultivating medium is between about 34 and about 38 degrees centigrade.

[CLAIM 9] The method according to claim 8, wherein the temperature is about 36 degrees centigrade.

[CLAIM 10] The method according to any one of claims 1-9, wherein said carbon source further comprises glucose.

[CLAIM 11] The method according to any one of claims 1-10, wherein said cultivating further comprises monitoring an OD of the cultivating medium such that when the OD reaches a designated level, said linear addition of a carbon source is initiated.

[CLAIM 12] The method according to claim 11, wherein said designated level is selected to achieve a higher volumetric production of cps.

[CLAIM 13] The method according to claim 11 or claim 12, wherein said designated level is between about 9.8 and about 10.2.

[CLAIM 14] The method according to claim 13, wherein said designated level is about 10.

[CLAIM 15] The method according to any one of claims 11-14, wherein said cultivating further comprises monitoring an OD of the cultivating

- medium such that when the OD reaches a first and second instantaneous addition level, two instantaneous additions of yeast extract are initiated prior to said linear addition of a carbon source.
- [CLAIM 16] The method according to claim 15, wherein said first and second instantaneous addition levels are selected to achieve a higher volumetric production of cps.
- [CLAIM 17] The method according to claim 16, wherein said first instantaneous addition level is between about 2.8 and about 3.2.
- [CLAIM 18] The method according to claim 17, wherein said first instantaneous addition level is about 3.0.
- [CLAIM 19] The method according to any one of claims 16-18, wherein said second instantaneous addition level is between about 4.3 and about 4.7. [CLAIM 20] The method according to claim 19, wherein said second
- instantaneous addition level is about 4.5.
- [CLAIM 21] The method according to any one of claims 1-20, wherein said cultivating medium is a defined medium comprising a phosphate source, a mineral source, a carbon source, a vitamin source, and an amino acid source to grow Streptococus, wherein said vitamin source consists of six or fewer vitamins selected from the following list of seven vitamins: biotin, niacinamide, calcium pantothenate, riboflavin, thiamine hydrochloride, pyridoxine hydrochloride and folic acid, wherein two of the vitamins have to be calcium pantothenate and niacinamide.
- [CLAIM 22] The method according to any one of claims 1-20, wherein said cultivating medium is a complex medium comprising a yeast extract, a phosphate source, a carbon source, a vitamin source, and optionally an amino acid source to grow Streptococcus, wherein said vitamin source consists of four or fewer vitamins selected from the following list of five vitamins: biotin, niacinamide, riboflavin, thiamine hydrochloride and pyridoxine hydrochloride, wherein one of the vitamins has to be biotin.
- [CLAIM 23] A cultivating medium comprising a Streptococcus strain, a phosphate source, a carbon source, a vitamin source, and an amino acid source to grow Streptococcus, wherein said vitamin source consists of six or fewer vitamins selected from the following list of seven vitamins: biotin, niacinamide, calcium pantothenate, riboflavin, thiamine hydrochloride, pyridoxine hydrochloride and folic acid, wherein two of the vitamins have to be calcium pantothenate and niacinamide.
- [CLAIM 24] The cultivating medium according to claim 23, wherein said Streptococcus is Streptococcus agalactiae.
- [CLAIM 25] The cultivating medium according to claim 23 or claim 24, wherein said phosphate source consists of K2HPO4, KH2PO4, Na2HPO4-H2O, NaH2PO4-H2O, or NaCl.
- [CLAIM 26] The cultivating medium according to any one of claims 23-25, wherein said carbon source is glucose.
- [CLAIM 27] The cultivating medium according to any one of claims 23-26, wherein said vitamin source consists of six or fewer from the following list of seven vitamins: biotin, niacinamide, calcium pantothenate, riboflavin, thiamine hydrochloride, pyridoxine hydrochloride, and folic acid, wherein two have to be calcium pantothenate and niacinamide.
- [CLAIM 28] The cultivating medium according to any one of claims 23-26, wherein said vitamin source consists of five or fewer from the following list of seven vitamins: biotin, niacinamide, calcium pantothenate, riboflavin, thiamine hydrochloride, pyridoxine hydrochloride, and folic acid, wherein two have to be calcium pantothenate and niacinamide.
- [CLAIM 29] The cultivating medium according to any one of claims 23-26, wherein said vitamin source consists of four or fewer from the following list of seven vitamins: biotin, niacinamide, calcium pantothenate, riboflavin, thiamine hydrochloride, pyridoxine hydrochloride, and folic acid, wherein two have to be calcium

pantothenate and niacinamide.

- [CLAIM 30] The cultivating medium according to any one of claims 23-26, wherein said vitamin source consists of three or fewer from the following list of seven vitamins: biotin, niacinamide, calcium pantothenate, riboflavin, thiamine hydrochloride, pyridoxine hydrochloride, and folic acid, wherein two have to be calcium pantothenate and niacinamide.
- [CLAIM 31] The cultivating medium according to any one of claims 23-26, wherein said vitamin source consists of calcium pantothenate and niacinamide.
- (CLAIM 32) The cultivating medium according to any one of claims 23-31, wherein said amino acid source consists of nineteen or fewer from the following list of nineteen amino acids: alanine, arginine, glutamine, glycine, histidine, isoleucine, leucine, lysine, methionine, phenylalanine, proline, serine, threonine, tryptophan, valine, aspartic acid, cysteine hydrochloride, glutamic acid, and tyrosine, wherein fifteen have to be arginine, glycine, histidine, isoleucine, leucine, lysine, methionine, phenylalanine, serine, threonine, tryptophan, valine, cysteine hydrochloride, glutamic acid, and tyrosine.
- (CLAIM 33) The cultivating medium according to any one of claims 23-31, wherein said amino acid source consists of eighteen or fewer from the following list of nineteen amino acids: alanine, arginine, glutamine, glycine, histidine, isoleucine, leucine, lysine, methionine, phenylalanine, proline, serine, threonine, tryptophan, valine, aspartic acid, cysteine hydrochloride, glutamic acid, and tyrosine, wherein fifteen have to be arginine, glycine, histidine, isoleucine, leucine, lysine, methionine, phenylalanine, serine, threonine, tryptophan, valine, cysteine hydrochloride, glutamic acid, and tyrosine.
- [CLAIM 34] The cultivating medium according to any one of claims 23-31, wherein said amino acid source consists of seventeen or fewer from the following list of nineteen amino acids: alanine, arginine, glutamine, glycine, histidine, isoleucine, leucine, lysine, methionine, phenylalanine, proline, serine, threonine, tryptophan, valine, aspartic acid, cysteine hydrochloride, glutamic acid, and tyrosine, wherein fifteen have to be arginine, glycine, histidine, isoleucine, leucine, lysine, methionine, phenylalanine, serine, threonine, tryptophan, valine, cysteine hydrochloride, glutamic acid, and tyrosine.
- [CLAIM 35] The cultivating medium according to any one of claims 23-31, wherein said amino acid source consists of sixteen or fewer from the following list of nineteen amino acids: alanine, arginine, glutamine, glycine, histidine, isoleucine, leucine, lysine, methionine, phenylalanine, proline, serine, threonine, tryptophan, valine, aspartic acid, cysteine hydrochloride, glutamic acid, and tyrosine, wherein fifteen have to be arginine, glycine, histidine, isoleucine, leucine, lysine, methionine, phenylalanine, serine, threonine, tryptophan, valine, cysteine hydrochloride, glutamic acid, and tyrosine.
- [CLAIM 36] The cultivating medium according to any one of claims 23-31, wherein said amino acid source consists of arginine, glycine, histidine, isoleucine, leucine, lysine, methionine, phenylalanine, serine, threonine, tryptophan, valine, cysteine hydrochloride, glutamic acid, and tyrosine.
- [CLAIN 37] A cultivating medium comprising a Streptococcus strain, a yeast extract, a phosphate source, a cateon source, a vitamin source, and optionally an amino acid source to grow Streptococcus, wherein said vitamin source consists of four or fewer vitamins selected from the following list of five vitamins: biotin, niacinamide, riboflavin, thiamine hydrochloride and pyridoxine hydrochloride, wherein one of the vitamins has to be biotin.
- [CLAIM 38] The cultivating medium according to claim 37, wherein said Streptococcus is Streptococcus agalactiae.
- [CLAIM 39] The cultivating medium according to claim 37 or claim 38, wherein said vitamin source consists of four or fewer vitamins selected

- from the following list of five vitamins: biotin, niacinamide, riboflavin, thiamine hydrochloride and pyridoxine hydrochloride, wherein one of the vitamins has to be biotin.
- [CLAIM 40] The cultivating medium according to claim 37 or claim 38, wherein said vitamin source consists of three or fewer vitamins selected from the following list of five vitamins: biotin, niacinamide, riboflavin, thiamine hydrochloride and pyridoxine hydrochloride, wherein one of the vitamins has to be biotin.
- [CLAIM 41] The cultivating medium according to claim 37 or claim 38, wherein said vitamin source consists of two or fewer vitamins selected from the following list of five vitamins: biotin, nlacinamide, riboflavin, thiamine hydrochloride and pyridoxine hydrochloride, wherein one of the vitamins has to be biotin.
- [CLAIM 42] The cultivating medium according to claim 37 or claim 38, wherein said vitamin source consists of biotin.
- [CLAIM 43] A method for purifying a capsular polysaccharide from Streptococcus agalactiae comprising a step of filtration using an adherent filter.
- [CLAIM 44] The method according to claim 43, wherein the method does not include a step of cationic detergent treatment to precipitate the capsular polysaccharide followed by a step of re-solubilization of the capsular polysaccharide.
- [CLAIM 45] The method according to claim 43 or claim 44, wherein the adherent filter is a protein adherent filter.
- [CLAIM 46] The method according to any of claims 43-45, wherein the adherent filter is a carbon filter.
- [CLAIM 47] The method according to any of claims 43-46, wherein the step of filtration using an adherent filter is preceded by the following steps: (i) alcoholic precipitation of contaminating proteins and/or nucleic acids; and (ii) diafiltration;
- [CLAIM 48] The method according to any of claims 43-47, wherein the step of filtration using an adherent filter is followed by the following steps: (iv) re-N-acetylation; (v) diafiltration.
- [CLAIM 49] A method for production of a purified capsular polysaccharide comprising: (a) providing a crude isolate containing a capsular polysaccharide; (b) removing an alcohol precipitate formed by contacting the crude isolate with an alcohol solution; (c) filtering to remove smaller molecular weight compounds while retaining the capsular polysaccharide; and (d) removing protein contaminants with a protein adherent filter to produce the purified capsular polysaccharide.
- [CLAIM 50] The method according to claim 49 further comprising step (e) re-N-acetylating the purified capsular polysaccharide.
- [CLAIM 51] The method according to claim 50 further comprising step (f) precipitating the purified capsular polysaccharide.
- [CLAIM 52] The method according to claim 51 further comprising step (g) formulating a vaccine with the capsular polysaccharide as a component.
- [CLAIM 53] The method according to any one of claims 49-52, wherein step (b) comprises addition of an alcohol solution to a concentration sufficient to precipitate nucleic acidcontaminants but not the capsular polysaccharide.
- [CLAIM 54] The method according to claim 53, wherein said alcohol solution comprises ethanol.
- [CLAIM 55] The method according to claim 53 or claim 54, wherein said alcohol solution further comprises CaCl2.
- [CLAIM 56] The method according to claim 54 or claim 55, wherein said alcohol solution is added to a concentration of between about 10 % and about 50% ethanol.
- [CLAIM 57] The method according to claim 56, wherein said alcohol solution is added to a concentration of about 30% ethanol.
- [CLAIM 58] The method according to any one of claims 49-57, wherein the protein adherent filter is an activated carbon filter.

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**FERMENTATIONSVERFAHREN ZUR KULTIVIERUNG VON STREPTOKOKKEN UND
    REINIGUNGSVERFAHREN ZUR GEWINNUNG VON CPS DARAUS
  FERMENTATION PROCESSES FOR CULTIVATING STREPTOCOCCI AND PURIFICATION
    PROCESSES FOR OBTAINING CPS THEREFROM
  PROCEDES DE FERMENTATION POUR CULTIVER DES STREPTOCOQUES ET PROCEDES DE
    PURIFICATION POUR OBTENIR DES CPS A PARTIR DE CEUX-CI**
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Language: EN
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Original IPC: A61K-39/04(B,I,H,EP,20060101,20090722,A,L)
   A61K-39/04(B,I,M,98,20060101,20090722,C)
    C12N-1/20(B, I, H, EP, 20060101, 20090722, A, F)
    C12N-1/20(B, I, M, 98, 20060101, 20090722, C)
    C12P-19/00(B, I, M, 98, 20060101, 20090722, C)
    C12P-19/04(B, I, H, EP, 20060101, 20090722, A, L)
Current IPC: A61K-39/04(B,I,H,EP,20060101,20090722,A,L)
    A61K-39/04(B, I, M, 98, 20060101, 20090722, C)
    C12N-1/20(B, I, H, EP, 20060101, 20090722, A, F)
    C12N-1/20(B, I, M, 98, 20060101, 20090722, C)
    C12P-19/00(B, I, M, 98, 20060101, 20090722, C)
    C12P-19/04(B, I, H, EP, 20060101, 20090722, A, L)
Current ECLA class: C08B-37/00P C12N-1/20 C12P-19/04
Current ECLA ICO class: K61K-39:00
Original Abstract: This invention is in the field of bacterial cultures,
    and specifically relates to the optimization of culture conditions to
    improve the production of bacterial capsular polysaccharides from
    Streptococcus strains in fed batch culture and to novel purification
    methods suitable for production scale purification of bacterial
    capsular polysaccharides from Streptococcus strains resulting in higher
    levels of purity than previously obtained for production scale.
Publication No. IN 201002275 P2 (Update 201072 E)
Publication Date: 20101008
**Fermentation processes for cultivating streptococci and purification
    processes for obtaining capsular polysaccharide of streptococci**
Assignee: NOVARTIS AG; CH (NOVS)
Inventor: CONSTANTINO P
  NORELLI F
  BERTI F
  CICALA C M
  BAZZOCCHI G
  FONTANI S
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OLIVIERI R
Language: EN
Application: IN 2010KN2275 A 20100622 (Local application)
  WO 2008IB3729 A 20081219 (PCT Application)
Priority: US 20078941 P 20071220
Original IPC: C12N-1/20(A) C12P-19/04(B)
Current IPC: C12N-1/20(A) C12P-19/04(B)
Japan
Publication No. JP 2011507501 W (Update 201118 E)
Publication Date: 20110310
Language: JA (120 pages)
Application: JP 2010538947 A 20081219 (Local application)
  WO 2008IB3729 A 20081219 (PCT Application)
Priority: US 20078941 P 20071220
  GB 200818453 A 20081008
Related Publication: WO 2009081276 A (Based on OPI patent )
Original IPC: A61K-39/00(B,I,H,JP,20060101,20110210,A,L)
    A61K-39/39(B, I, H, JP, 20060101, 20110210, A, L)
    A61P-31/04(B, I, H, JP, 20060101, 20110210, A, L)
    C12N-1/20(B, I, H, JP, 20060101, 20110210, A, F)
    C12P-19/04(B, I, H, JP, 20060101, 20110210, A, L)
Current IPC: A61K-39/00(B,I,H,JP,20060101,20110210,A,L)
    A61K-39/39(B, I, H, JP, 20060101, 20110210, A, L)
    A61P-31/04(B, I, H, JP, 20060101, 20110210, A, L)
    C12N-1/20(B, I, H, JP, 20060101, 20110210, A, F)
    C12P-19/04(B, I, H, JP, 20060101, 20110210, A, L)
Current JP FI-Terms: C12N-1/20 A (main, A) A61K-39/00 G (secondary, B)
    A61K-39/39 (secondary, B) A61P-31/04 (secondary, B) C12P-19/04 C
    (secondary, B) C12N-1/20 A (linked, C) C12P-19/04 C (linked, C)
    C12R-1:46 (linked, C)
Current JP F-Terms: 4B064 4B065 4C085 4C085AA03 4B065AA49X 4B065AC20
    4B064AF17 4C085BA14 4C085BA38 4B065BB02 4B065BB06 4B065BB12 4B065BB15
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    4B064CE20 4B064DA01 4C085DD21 4C085DD24 4C085DD25 4C085DD26 4C085DD31
    4C085DD37 4C085DD41 4C085EE01 4C085GG01 4C085GG08 4C085GG10
United States
Publication No. US 20100272755 A1 (Update 201071 E)
Publication Date: 20101028
**FERMENTATION PROCESSES FOR CULTIVATING STREPTOCOCCI AND PURIFICATION
    PROCESSES FOR OBTAINING CPS THEREFROM**
Assignee: NOVARTIS VACCINES AND DIAGNOSTICS SRL, Siena, IT (NOVS)
  Costantino, Paolo, Colle Val d'Elsa, IT Residence: IT
  Norelli, Francesco, Siena, IT Residence: IT
  Berti, Francesco, Colle Val d'Elsa, IT Residence: IT
  Olivieri, Roberto, Siena, IT Residence: IT
  Bazzocchi, Giulia, Vicenza, IT Residence: IT
  Cicala, Concetta Maria, Castellina Scalo, IT Residence: IT
  Fontani, Silvia, Siena, IT Residence: IT
Inventor: COSTANTINO P, IT
 Norelli, Francesco, Siena, IT Residence: IT
  Berti, Francesco, Colle Val d'Elsa, IT Residence: IT
 Olivieri, Roberto, Siena, IT Residence: IT
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  Cicala, Concetta Maria, Castellina Scalo, IT Residence: IT
  Fontani, Silvia, Siena, IT Residence: IT
Agent: NOVARTIS VACCINES AND DIAGNOSTICS INC., INTELLECTUAL PROPERTY-
   X100B, P.O. BOX 8097, Emeryville, CA, US
Language: EN
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Application: US 2010747914 A 20100613 (Local application)
  WO 2008IB3729 A 20081219 (PCT Application)
  US 20078941 P 20071220 (Related to Provisional)
Priority: GB 200818453 A 20081008
Original IPC: A61K-39/09(B,I,H,US,20060101,20101028,A,F)
    A61K-39/09(B, I, M, 98, 20060101, 20101028, C)
    C07H-1/00(B, I, H, US, 20060101, 20101028, A, L)
    C07H-1/00(B, I, M, 98, 20060101, 20101028, C)
    C12N-1/20(B, I, H, US, 20060101, 20101028, A, L)
    C12N-1/20(B,I,M,98,20060101,20101028,C)
    C12P-19/00(B, I, M, 98, 20060101, 20101028, C)
    C12P-19/04(B, I, H, US, 20060101, 20101028, A, L)
    C12Q-3/00(B, I, H, US, 20060101, 20101028, A, L)
    C12Q-3/00(B, I, M, 98, 20060101, 20101028, C)
Current IPC: A61K-39/09(B,I,H,US,20060101,20101028,A,F)
    A61K-39/09(B, I, M, 98, 20060101, 20101028, C)
    C07H-1/00(B, I, H, US, 20060101, 20101028, A, L)
    C07H-1/00(B, I, M, 98, 20060101, 20101028, C)
    C12N-1/20(B, I, H, US, 20060101, 20101028, A, L)
    C12N-1/20(B, I, M, 98, 20060101, 20101028, C)
    C12P-19/00(B,I,M,98,20060101,20101028,C)
    C12P-19/04(B, I, H, US, 20060101, 20101028, A, L)
    C120-3/00(B, I, H, US, 20060101, 20101028, A, L)
    C12Q-3/00(B, I, M, 98, 20060101, 20101028, C)
Current ECLA class: C08B-37/00P C12N-1/20 C12P-19/04
Current ECLA ICO class: K61K-39:00
Current US Class (main): 424-244100
Current US Class (secondary): 435-003000 435-101000 435-253400 536-123100
Original US Class (main): 424244.1
Original US Class (secondary): 4353 435101 435253.4 536123.1
Original Abstract: This invention is in the field of bacterial cultures
    and specifically relates to the optimization of culture conditions to
    improve the production of bacterial capsular polysaccharides from
    ~Streptococcus~ strains in fed batch culture and to novel purification
    methods suitable for production scale purification of bacterial
    capsular polysaccharides from ~Streptococcus~ strains resulting in
    higher levels of purity than previously obtained for production scale.
Claim:
 1.
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1. A method for cultivating ~Streptococcus~ for production of capsular polysaccharides (cps) on a manufacturing scale, wherein said method comprises (a) providing an inoculum of a strain of ~Streptococcus~ expressing the cps, and (b) cultivating the strain by fermentation, wherein said cultivating comprises a linear addition of a carbon source to a cultivating medium, and does not use an algorithm to control the cultivating by monitoring a pH of the cultivating medium.

WIPO

Publication No. WO 2009081276 A2 (Update 200946 B)

Publication Date: 20090702

**FERMENTATION PROCESSES FOR CULTIVATING STREPTOCOCCI AND PURIFICATION PROCESSES FOR OBTAINING CPS THEREFROM PROCEDES DE FERMENTATION POUR CULTIVER DES STREPTOCOOUES ET PROCEDES DE

PURIFICATION POUR OBTENIR DES CPS A PARTIR DE CEUX-CI** Assignee: ~(except US)~ NOVARTIS AG, Lichtstrasse 35, CH-4056 Basel, CH

Residence: CH Nationality: CH (NOVS) ~(only US)~ COSTANTINO, Paolo, Novartis Vaccines, Via Fiorentina 1,

I-53100 Siena, IT Residence: IT Nationality: IT ~(only US)~ NORELLI, Francesco, Novartis Vaccines, Via Fiorentina 1,

I-53100 Siena, IT Residence: IT Nationality: IT

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  ~(only US)~ CICALA, Concetta, Maria, Novartis Vaccines, Via Fiorentina 1,
    I-53100 Siena, IT Residence: IT Nationality: IT
  ~(only US)~ BAZZOCCHI, Giulia, Novartis Vaccines, Via Fiorentina 1,
    I-53100 Siena, IT Residence: IT Nationality: IT
  ~(only US)~ FONTANI, Silvia, Novartis Vaccines, Via Fiorentina 1, I-53100
    Siena, IT Residence: IT Nationality: IT
  ~(only US)~ OLIVIERI, Roberto, Novartis Vaccines, Via Fiorentina 1,
    I-53100 Siena, IT Residence: IT Nationality: IT
Inventor: BAZZOCCHI, Giulia, Novartis Vaccines, Via Fiorentina 1, I-53100
    Siena, IT Residence: IT Nationality: IT
  BERTI, Francesco, Novartis Vaccines, Via Fiorentina 1, I-53100 Siena, IT
    Residence: IT Nationality: IT
  CICALA, Concetta, Maria, Novartis Vaccines, Via Fiorentina 1, I-53100
    Siena, IT Residence: IT Nationality: IT
  FONTANI, Silvia, Novartis Vaccines, Via Fiorentina 1, I-53100 Siena, IT
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  NORELLI, Francesco, Novartis Vaccines, Via Fiorentina 1, I-53100 Siena,
    IT Residence: IT Nationality: IT
  OLIVIERI, Roberto, Novartis Vaccines, Via Fiorentina 1, I-53100 Siena, IT
    Residence: IT Nationality: IT
  COSTANTINO, Paolo, Novartis Vaccines, Via Fiorentina 1, I-53100 Siena, IT
Agent: MARSHALL, Cameron, John et al., Carpmaels Ransford, 43-45
    Bloomsbury Square, London WC1A 2RA, GB
Language: EN (159 pages, 30 drawings)
Application: WO 2008IB3729 A 20081219 (Local application)
Priority: US 20078941 P 20071220
  GB 200818453 A 20081008
Designated States: (National Confirmed) AE AG AL AM AO AT AU AZ BA BB BG BH
    BR BW BY BZ CA CH CN CO CR CU CZ DE DK DM DO DZ EC EE EG ES FI GB GD GE
    GH GM GT HN HR HU ID IL IN IS JP KE KG KM KN KP KR KZ LA LC LK LR LS LT
    LU LY MA MD ME MG MK MN MW MX MY MZ NA NG NI NO NZ OM PG PH PL PT RO RS
    RU SC SD SE SG SK SL SM ST SV SY TJ TM TN TR TT TZ UA UG US UZ VC VN ZA
    ZM ZW
  (Regional Confirmed) AT BE BG CH CY CZ DE DK EE ES FI FR GB GR HR HU IE
    IS IT LT LU LV MC MT NL NO PL PT RO SE SI SK TR OA BW GH GM KE LS MW MZ
    NA SD SL SZ TZ UG ZM ZW EA
Original IPC: A61K-39/04(B,I,H,EP,20060101,A,L)
    A61K-39/04(B,I,M,98,20060101,C) C12N-1/20(B,I,H,EP,20060101,A,F)
    C12N-1/20(B,I,M,98,20060101,C) C12P-19/00(B,I,M,98,20060101,C)
    C12P-19/04(B, I, H, EP, 20060101, A, L)
Current IPC: A61K-39/04(B,I,H,EP,20060101,20090702,A,L)
    A61K-39/04(B, I, H, EP, 20090101, 20090702, C, L)
    C12N-1/20(B, I, H, EP, 20060101, 20090702, A, F)
    C12N-1/20(B, I, H, EP, 20090101, 20090702, C, F)
    C12P-19/00(B,I,H,EP,20090101,20090702,C,L)
    C12P-19/04(B, I, H, EP, 20060101, 20090702, A, L)
Current ECLA class: C08B-37/00P C12N-1/20 C12P-19/04
Current ECLA ICO class: K61K-39:00
Original Abstract: This invention is in the field of bacterial cultures,
    and specifically relates to the optimization of culture conditions to
    improve the production of bacterial capsular polysaccharides from
    Streptococcus strains in fed batch culture and to novel purification
    methods suitable for production scale purification of bacterial
    capsular polysaccharides from Streptococcus strains resulting in higher
    levels of purity than previously obtained for production scale.
   La presente invention concerne le domaine des cultures bacteriennes, et
    concerne specifiquement l'optimisation de conditions de culture pour
    ameliorer la production de polysaccharides capsulaires bacteriens a
    partir de souches de Streptococcus (CPS) en culture a ecoulement
   discontinu. L'invention concerne egalement des nouveaux procedes de
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purification adaptes pour la purification a l'echelle industrielle de
    polysaccharides capsulaires bacteriens a partir de souches de
    Streptococcus resultant en des niveaux plus eleves de purete que ceux
    precedemment obtenus a l'echelle industrielle.
Publication No. WO 2009081276 A3 (Update 200958 E)
Publication Date: 20090903
**FERMENTATION PROCESSES FOR CULTIVATING STREPTOCOCCI AND PURIFICATION
    PROCESSES FOR OBTAINING CPS THEREFROM**
Assignee: NOVARTIS AG: CH (NOVS)
Inventor: BAZZOCCHI G, IT
  BERTI F, IT
  CICALA C M, IT
  COSTANTINO P, IT
  FONTANI S, IT
 NORELLI F. IT
  OLIVIERI R, IT
Language: EN
Application: WO 2008IB3729 A 20081219 (Local application)
Priority: US 20078941 P 20071220
  GB 200818453 A 20081008
Designated States: (National Confirmed) AE AG AL AM AO AT AU AZ BA BB BG BH
   BR BW BY BZ CA CH CN CO CR CU CZ DE DK DM DO DZ EC EE EG ES FI GB GD GE
    GH GM GT HN HR HU ID IL IN IS JP KE KG KM KN KP KR KZ LA LC LK LR LS LT
    LU LY MA MD ME MG MK MN MW MX MY MZ NA NG NI NO NZ OM PG PH PL PT RO RS
    RU SC SD SE SG SK SL SM ST SV SY TJ TM TN TR TT TZ UA UG US UZ VC VN ZA
    ZM ZW
  (Regional Confirmed) AT BE BG CH CY CZ DE DK EE ES FI FR GB GR HR HU IE
    IS IT LT LU LV MC MT NL NO PL PT RO SE SI SK TR OA BW GH GM KE LS MW MZ
    NA SD SL SZ TZ UG ZM ZW EA
Original IPC: A61K-39/04(B, I, H, EP, 20060101, A, L)
    A61K-39/04(B,I,M,98,20060101,C) C12N-1/20(B,I,H,EP,20060101,A,F)
    C12N-1/20(B,I,M,98,20060101,C) C12P-19/00(B,I,M,98,20060101,C)
    C12P-19/04(B, I, H, EP, 20060101, A, L)
Current IPC: A61K-39/04(B,I,H,EP,20060101,20090903,A,L)
    A61K-39/04(B, I, H, EP, 20090101, 20090903, C, L)
    C12N-1/20(B, I, H, EP, 20060101, 20090903, A, F)
    C12N-1/20(B, I, H, EP, 20090101, 20090903, C, F)
    C12P-19/00(B, I, H, EP, 20090101, 20090903, C, L)
    C12P-19/04(B, I, H, EP, 20060101, 20090903, A, L)
Current ECLA class: C08B-37/00P C12N-1/20 C12P-19/04
Current ECLA ICO class: K61K-39:00
Original Abstract: This invention is in the field of bacterial cultures,
    and specifically relates to the optimization of culture conditions to
    improve the production of bacterial capsular polysaccharides from
    Streptococcus strains in fed batch culture and to novel purification
    methods suitable for production scale purification of bacterial
    capsular polysaccharides from Streptococcus strains resulting in higher
    levels of purity than previously obtained for production scale.
 10/7/7
            (Item 7 from file: 351)
DIALOG(R)File 351:Derwent WPI
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0018714931
WPI ACC NO: 2009-E02825/200914
Purifying saccharide antigen-carrier protein conjugates from a mixture, comprises contacting the mixture with hydroxyapatite and collecting the free saccharide antigen-carrier protein conjugates
Patent Assignee: NOVARTIS AG (NOVS); NOVARTIS VACCINES&DIAGNOSTICS INC (NOVS)
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Inventor: AVERANI G; BELLUCCI C; BERTI F; BIGIO M; NORELLI F; AVERANI G, CH ; BELLUCCI C, CH; BERTI F, CH; BIGIO M, CH; NORELLI F, CH

Patent Family (7 patents, 123 countries)

Application Patent Number Kind Date Number Kind Update Date WO 2009010877 A2 20090122 WO 2008IB2690 A 20080717 200914 WO 2009010877 A3 20091119 WO 2008IB2690 A 20080717 200976 AU 2008277353 A1 20090122 AU 2008277353 A 20080717 201016 Ε EP 2180901 A2 20100505 EP 2008826340 A 20080717 201030 E WO 2008IB2690 A 20080717 CA 2693936 A1 20090122 CA 2693936 A 20080717 201031 E WO 2008IB2690 A 20080717 CA 2693936 A 20100118 CN 101795713 20100804 CN 200880106077 A 20080717 201058 WO 2008IB2690 A 20080717 US 20100239600 20100923 WO 2008IB2690 A 20080717 A1 201062 E US 2010669464 A 20100609

Priority Applications (no., kind, date): GB 200713880 A 20070717

Patent Details

Number Kind Lan Pa Dwa Filina Notes

WO 2009010877 A2 EN 54 National Designated States, Original: AE AG AL AM AO AT AU AZ BA BB BG BH BR BW BY BZ CA CH CN CO CR CU CZ DE DK DM DO DZ EC EE EG ES FI GB GD GE GH GM GT HN HR HU ID IL IN IS JP KE KG KM KN KP KR KZ LA LC LK LR LS LT LU LY MA MD ME MG MK MN MW MX MY MZ NA NG NI NO NZ OM PG PH PL PT RO RS RU SC SD SE SG SK SL SM ST SV SY TJ TM TN TR TT TZ UA UG US UZ VC VN ZA ZM ZW

Regional Designated States, Original: AT BE BG CH CY CZ DE DK EE ES FI FR GB GR HR HU IE IS IT LT LU LV MC MT NL NO PL PT RO SE SI SK TR OA BW GH GM KE LS MW MZ NA SD SL SZ TZ UG ZM ZW EA

WO 2009010877

A3 EN National Designated States, Original: AE AG AL AM AO AT AU AZ BA BB BG BH BR BW BY BZ CA CH CN CO CR CU CZ DE DK DM DO DZ EC EE EG ES FI GB GD GE GH GM GT HN HR HU ID IL IN IS JP KE KG KM KN KP KR KZ LA LC LK LR LS LT LU LY MA MD ME MG MK MN MW MX MY MZ NA NG NI NO NZ OM PG PH PL PT RO RS RU SC SD SE SG SK SL SM ST SV SY TJ TM TN TR TT TZ UA UG US UZ VC VN ZA

Regional Designated States, Original: AT BE BG CH CY CZ DE DK EE ES FI FR GB GR HR HU IE IS IT LT LU LV MC MT NL NO PL PT RO SE SI SK TR OA BW GH GM KE LS MW MZ NA SD SL SZ TZ UG ZM ZW EA

AU 2008277353 A1 EN Based on OPI patent

WO 2009010877 EP 2180901 A2 EN PCT Application WO 2008IB2690 Based on OPI patent WO 2009010877

Regional Designated States, Original: AT BE BG CH CY CZ DE DK EE ES FI FR GB GR HR HU IE IS IT LI LT LU LV MC MT NL NO PL PT RO SE SI SK TR

CA 2693936 A1 EN PCT Application WO 2008IB2690 PCT national entry CA 2693936 Based on OPI patent WO 2009010877

PCT Application WO 2008IB2690 CN 101795713 Based on OPI patent WO 2009010877 US 20100239600 A1 PCT Application WO 2008IB2690

Alerting Abstract WO A2

NOVELTY - Purifying saccharide antigen-carrier protein conjugates from a mixture, comprises contacting the mixture with hydroxyapatite and collecting the free saccharide antigen-carrier protein conjugates. DESCRIPTION - INDEPENDENT CLAIMS are:

1.a method of preparing a pharmaceutical composition, comprising the method above, and mixing the saccharide antigen-carrier protein

2.a pharmaceutical composition prepared by the latter method, for use (i) in therapy, (ii) for raising an immune response or (iii) as a vaccine.

ACTIVITY - Antibacterial. No biological data given. MECHANISM OF ACTION - Vaccine; Protease-Inhibitor.

USE - The methods and composition are useful for purifying saccharide antigen-carrier protein conjugates from a mixture; for preparing a pharmaceutical composition; for use in therapy, for raising an immune response or as a vaccine; and for manufacturing a medicament for raising the immune response, or treating a bacterial infection (all claimed).

Technology Focus

BIOTECHNOLOGY - Preferred Method: In purifying saccharide antigen-carrier protein conjugates from the mixture, the mixture comprises free carrier protein and saccharide antigen-carrier protein conjugates. The mixture also comprises other contaminant proteins. The carrier protein is selected from tetanus toxoid, diphtheria toxoid, derivatives, ~****Neisseria**** ****meningitidis**** ~ outer membrane proteins, synthetic proteins, heat shock proteins, pertussis proteins, cytokines, lymphokines, hormones, growth factors, poly-epitope carriers, protein D of ~Haemophilus influenzae , pneumolysin, pneumococcal surface protein PspA, iron uptake proteins, toxin A or B from ~C. difficile ~ and/or a polyepitope carrier such as N19. The carrier protein is tetanus toxoid, diphtheria toxoid or derivatives. The carrier protein is CRM197. The saccharide antigen has a molecular weight of 5 kDa or 50 kDa or more. Also, the saccharide antigen is a bacterial ****capsular**** saccharide. Also, the saccharide antigen is glycosylated. Furthermore, the saccharide antigen is from ~N. ****meningitidis**** ~ , ~Streptococcus pneumoniae ~ , ~Streptococcus agalactiae ~ , ~H. influenzae ~ , ~Pseudomonas aeruginosa ~ , ~Staphylococcus aureus ~ , ~E. faecalis ~ , ~E. faecium ~ , ~Y. enterocolitica ~ , ~V. cholerae ~ or ~S. typhi ~ . The carrier protein is conjugated to saccharide antigens from more than one bacterial species. The saccharide antigen is also conjugated to the carrier protein by a linker. Moreover, the method is carried out at pH 6.5-7.5, preferably at pH 7.2. Also, the method is carried out at a phosphate concentration of 50 mM or less. The hydroxyapatite is in the form of a gel, where the hydroxyapatite has a particle size of 40 mu m or more, and a dynamic binding capacity of more than 10 mg lysozyme/g. Preparing the pharmaceutical composition also comprises mixing the product with an adjuvant.

Title Terms/Index Terms/Additional Words: PURIFICATION; SACCHARIDE; ANTIGEN ; CARRY; PROTEIN; CONJUGATE; MIXTURE; COMPRISE; CONTACT; COLLECT; FREE

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Class Codes
International Classification (+ Attributes)
IRC + Level Value Position Status Version
A61K-0039/09 A I L B 20060101
A61K-0039/385 A I F B 20060101
A61K-0031/04 A I F B 20060101
A61P-0031/04 A I L B 20060101
A61P-0031/04 A I L B 20060101
A61P-0031/00 A I L B 20060101
B01D-0015/00 A I L B 20060101
A61K-0039/09 C I L B 20060101
A61K-0039/09 C I L B 20090101
A61K-0039/385 C I B 20060101
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A61K-0047/48 C I F B 20090101

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A61P-0031/00 C I L B 20090101
  A61P-0031/00 C I
                          B 20060101
  A61P-0037/00 C I
                          B 20060101
  B01D-0015/00 C I L B 20090101
  B01J-0020/04 C I L B 20090101
  C07K-0001/00 C I
                         B 20060101
                     20060101
  C12N S
ECLA: A61K-039/09A, A61K-047/48R2, A61K-047/48R2D, A61K-047/48R2F,
  A61K-047/48R2V, C07K-001/16
ICO: K61K-039:60P10
US Classification, Current Main: 424-193100; Secondary: 530-351000,
530-399000, 530-413000
US Classification, Issued: 424193.1, 530413, 530351, 530399
File Seament: CPI
DWPI Class: B04; D16
Manual Codes (CPI/A-M): B04-B04C1; B04-H01; B04-H06; B04-J01; B04-N03;
  B04-N09; B05-B02A3; B11-B03; B14-A01; B14-D07C; B14-G01; B14-S11; D05-H07
  ; D05-H13
Original Publication Data by Authority
Anstralia
Publication No. AU 2008277353 A1 (Update 201016 E)
Publication Date: 20090122
Assignee: NOVARTIS AG (NOVS)
Inventor: AVERANI G
  BELLUCCI C
  BERTI F
  BIGIO M
  NORELLI F
Language: EN
Application: AU 2008277353 A 20080717 (Local application)
Priority: GB 200713880 A 20070717
Related Publication: WO 2009010877 A (Based on OPI patent )
Original IPC: A61K-47/48(B,I,H,EP,20060101,20091216,A,F)
    A61K-39/09(B, I, H, EP, 20060101, 20091216, A, L)
    A61P-31/04(B, I, H, EP, 20060101, 20091216, A, L)
    B01D-15/00(B, I, H, EP, 20060101, 20091216, A, L)
    B01J-20/04(B,I,H,EP,20060101,20091216,A,L)
    A61K-39/09(B, I, H, EP, 20090101, 20091216, C, L)
    A61K-47/48(B, I, H, EP, 20090101, 20091216, C, F)
    A61P-31/00(B, I, H, EP, 20090101, 20091216, C, L)
    B01D-15/00(B, I, H, EP, 20090101, 20091216, C, L)
    B01J-20/04(B,I,H,EP,20090101,20091216,C,L)
Current IPC: A61K-39/09(B,I,H,EP,20060101,20091216,A,L)
    A61K-39/09(B,I,H,EP,20090101,20091216,C,L)
    A61K-47/48(B, I, H, EP, 20060101, 20091216, A, F)
    A61K-47/48(B, I, H, EP, 20090101, 20091216, C, F)
    A61P-31/00(B, I, H, EP, 20090101, 20091216, C, L)
    A61P-31/04(B, I, H, EP, 20060101, 20091216, A, L)
    B01D-15/00(B, I, H, EP, 20060101, 20091216, A, L)
    B01D-15/00(B, I, H, EP, 20090101, 20091216, C, L)
    B01J-20/04(B, I, H, EP, 20060101, 20091216, A, L)
    B01J-20/04(B, I, H, EP, 20090101, 20091216, C, L)
Current ECLA class: A61K-39/09A A61K-47/48R2 A61K-47/48R2D A61K-47/48R2F
    A61K-47/48R2V C07K-1/16
Current ECLA ICO class: K61K-39:60P10
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Publication No. CA 2693936 A1 (Update 201031 E)

```
Publication Date: 20090122
Assignee: NOVARTIS AG; CH (NOVS)
Inventor: AVERANI G, IT
  BELLUCCI C, IT
  BERTI F, IT
  BIGIO M. IT
 NORELLI F, IT
Language: EN
Application: CA 2693936 A 20080717 (Local application)
  WO 2008IB2690 A 20080717 (PCT Application)
  CA 2693936 A 20100118 (PCT national entry)
Priority: GB 200713880 A 20070717
Related Publication: WO 2009010877 A (Based on OPI patent )
Original IPC: A61K-39/09(B,I,H,EP,20060101,20100319,A,L)
    A61K-39/09(B, I, M, 98, 20060101, 20100319, C)
    A61K-47/48(B, I, H, EP, 20060101, 20100319, A, F)
    A61K-47/48(B, I, M, 98, 20060101, 20100319, C)
    A61P-31/00(B, I, M, 98, 20060101, 20100319, C)
    A61P-31/04(B, I, H, EP, 20060101, 20100319, A, L)
    B01D-15/00(B, I, H, EP, 20060101, 20100319, A, L)
    B01D-15/00(B, I, M, 98, 20060101, 20100319, C)
    B01J-20/04(B, I, H, EP, 20060101, 20100319, A, L)
    B01J-20/04(B,I,M,98,20060101,20100319,C)
Current IPC: A61K-39/09(B.I.H.EP.20060101.20100319.A.L)
    A61K-39/09(B, I, H, EP, 20100101, 20100319, C, L)
    A61K-47/48(B, I, H, EP, 20060101, 20100319, A, F)
    A61K-47/48(B, I, H, EP, 20100101, 20100319, C, F)
    A61P-31/00(B, I, H, EP, 20100101, 20100319, C, L)
    A61P-31/04(B, I, H, EP, 20060101, 20100319, A, L)
    B01D-15/00(B, I, H, EP, 20060101, 20100319, A, L)
    B01D-15/00(B, I, H, EP, 20100101, 20100319, C, L)
    B01J-20/04(B, I, H, EP, 20060101, 20100319, A, L)
    B01J-20/04(B, I, H, EP, 20100101, 20100319, C, L)
Current ECLA class: A61K-39/09A A61K-47/48R2 A61K-47/48R2D A61K-47/48R2F
    A61K-47/48R2V C07K-1/16
Current ECLA ICO class: K61K-39:60P10
Publication No. CN 101795713 A (Update 201058 E)
Publication Date: 20100804
**Conjugate purification**
Assignee: NOVARTIS AG: CH (NOVS)
Inventor: BERTI F. CH
  BERTI FRANCESCO, CH
  BIGIO M, CH
  BIGIO MASSIMO, CH
  AVERANI G, CH
  AVERANI GIOVANNI, CH
  NORELLI F. CH
  NORELLI FRANCESCO, CH
  BELLUCCI C, CH
  BELLUCCI CINZIA, CH
Language: ZH
Application: CN 200880106077 A 20080717 (Local application)
  WO 2008IB2690 A 20080717 (PCT Application)
Priority: GB 200713880 A 20070717
Related Publication: WO 2009010877 A (Based on OPI patent )
Original IPC: A61K-39/09(I,CN,20060101,A,L) A61K-39/09(I,M,98,20060101,C)
    A61K-47/48(I,CN,20060101,A,F) A61K-47/48(I,M,98,20060101,C)
    A61P-31/00(I,M,98,20060101,C) A61P-31/04(I,CN,20060101,A,L)
    B01D-15/00(I,CN,20060101,A,L) B01D-15/00(I,M,98,20060101,C)
    B01J-20/04(I,CN,20060101,A,L) B01J-20/04(I,M,98,20060101,C)
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Current IPC: A61K-39/09(B,I,H,CN,20060101,20100825,A,L)
A61K-39/09(B,I,H,CN,20100101,20100825,G,L)
A61K-47/48(B,I,H,CN,20060101,20100825,G,F)
A61K-47/48(B,I,H,CN,20100101,20100825,G,F)
A61F-31/09(B,I,H,CN,20100101,20100825,G,L)
A61F-31/09(B,I,H,CN,20100101,20100825,A,L)
B01D-15/00(B,I,H,CN,20060101,20100825,A,L)
B01D-15/00(B,I,H,CN,20060101,20100825,A,L)
B01D-15/00(B,I,H,CN,20060101,20100825,A,L)
B01D-15/00(B,I,H,CN,20060101,20100825,A,L)
```

- B01J-20/04(B,I,H,CN,20100101,20100825,C,L)
 Original Abstract: This application relates to methods for the purification of saccharide antigen-carrier protein conjugates. In particular, the invention provides a method for purifying saccharide antigen-carrier protein conjugates from free carrier protein, such as CRN197, using hydroxyapatite. The invention further relates to methods of preparing vaccines, using this method.
- Claim: [CLAIM 1] A method of purifying saccharide antigen-carrier protein conjugates from a mixture, comprising contacting said mixture with hydroxyapatite and collecting the free saccharide antigen-carrier protein conjugates.
 - [CLAIM 2] The method according to claim 1, wherein said mixture comprises free carrier protein and saccharide antigen-carrier protein conjugates. [CLAIM 3] The method according to claim 2, wherein the mixture further comprises other contaminant proteins.
 - [CLAIM 4] The method any one of foresaid claims, wherein the carrier protein is selected from tetanus toxoid, diphtheria toxoid, derivatives thereof, N. meningitidis outer membrane proteins, synthetic proteins, heat shock proteins, pertussis proteins, cytokines, lymphokines, hormones, growth factors, poly-epitope carriers, protein D of H. influenzae, pneumolysin, pneumococcal surface protein PspA, iron uptake proteins, toxin A or B from C. difficile and/or a poly epitope carrier such as N19.
 - [CLAIM 5] The method of any one according to claims 1-3, wherein said carrier protein is tetanus toxoid, diphtheria toxoid or derivatives thereof
 - [CLAIM 6] The method according to claim 5, wherein said carrier protein is CRM197.
 - [CLAIM 7] The method any one of foresaid claims, wherein the saccharide antigen has a molecular weight of $50 \mathrm{kDa}$ or more.
 - [CLAIM 8] The method according to claim 7, wherein the saccharide antigen has a molecular weight of 5kDa or more.
 - [CLAIM 9] The method any one of foresaid claims, wherein the saccharide antigen is a bacterial capsular saccharide.
 - [CLAIM 10] The method any one of foresaid claims, wherein the saccharide antigen is glycosylated.
 [CLAIM 11] The method any one of foresaid claims, wherein the saccharide
 - antigen is from N. meningitidis, S.pneumoniae, S.agalactiae, E[Lambda]nfluenzae, P. aeruginosa, S. aureus, E.faecalis, E.faechim, Y.enterocolitica, V.cholerae or S. typhi.
 - [CLAIM 12] The method any one of foresaid claims, wherein the earner protein is conjugated to saccharide antigens from more than one
 - bacterial species. [CLAIM 13] The method any one of foresaid claims, wherein the saccharide
 - antigen is conjugated to the carrier protein by a linker. [CLAIM 14] The method any one of foresaid claims, wherein said method is carried out at pH6.5-pH7.5.
 - [CLAIM 15] The method any one of foresaid claims, wherein said method is carried out at pH7.2.
 - [CLAIM 16] The method any one of foresaid claims, wherein said method is carried out at a phosphate concentration of 50mM or less.
 - [CLAIM 17] The method any one of foresaid claims, wherein said hydroxy apatite is in the form of a gel.

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[CLAIM 18] The method any one of foresaid claims, wherein said
    hydroxyapatite has a particle size of 40um or more.
  [CLAIM 19] The method any one of foresaid claims, wherein said
    hydroxyapatite has a dynamic binding capacity of more than 10mg
    lysozyme per gram.
  [CLAIM 20] A method of preparing a pharmaceutical composition, comprising
    the method any one of foresaid claims, and further comprising step iii)
    mixing said saccharide antigen-carrier protein conjugates obtained in
    step ii) with a pharmaceutically acceptable diluent or carrier
  [CLAIM 21] The method according to claim 20, further comprising step (iv)
    mixing the product of step (iii) with an adjuvant.
  [CLAIM 22] A pharmaceutical composition prepared by the method according
    to claim 20 or claim 21, for use (i) in therapy, (ii) for raising an
    immune response or (iii) as a vaccine.
  [CLAIM 23] Use of a pharmaceutical composition prepared by the method
    according to claim 20 or claim 21 in the manufacture of a medicament
    for (i) raising an immune response, or (ii) treating a bacterial
    infection.
Publication No. EP 2180901 A2 (Update 201030 E)
Publication Date: 20100505
**KONJUGATREINIGUNG
  CONJUGATE PURIFICATION
  PURIFICATION DE CONJUGUES**
Assignee: Novartis AG, Lichtstrasse 35, 4056 Basel, CH (NOVS)
Inventor: AVERANI, Giovanni, Novartis Vaccines, Via Fiorentina 1, I-53100
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  BELLUCCI, Cinzia, Novartis Vaccines, Via Fiorentina 1, I-53100 Siena, IT
  BERTI, Francesco, Novartis Vaccines, Via Fiorentina 1, I-53100 Siena, IT
  BIGIO, Massimo, Novartis Vaccines, Via Fiorentina 1, I-53100 Siena, IT
  NORELLI, Francesco, Novartis Vaccines, Via Fiorentina 1, I-53100 Siena,
Agent: Marshall, Cameron John, Carpmaels Ransford, 43-45 Bloomsbury
    Square, London WC1A 2RA, GB
Language: EN
Application: EP 2008826340 A 20080717 (Local application)
  WO 2008IB2690 A 20080717 (PCT Application)
Priority: GB 200713880 A 20070717
Related Publication: WO 2009010877 A (Based on OPI patent )
Designated States: (Regional Original) AT BE BG CH CY CZ DE DK EE ES FI FR
    GB GR HR HU IE IS IT LI LT LU LV MC MT NL NO PL PT RO SE SI SK TR
Original IPC: A61K-39/09(B,I,H,EP,20060101,20100224,A,L)
    A61K-39/09(B, I, M, 98, 20060101, 20100224, C)
    A61K-47/48(B, I, H, EP, 20060101, 20100224, A, F)
    A61K-47/48(B, I, M, 98, 20060101, 20100224, C)
    A61P-31/00(B, I, M, 98, 20060101, 20100224, C)
    A61P-31/04(B, I, H, EP, 20060101, 20100224, A, L)
    B01D-15/00(B, I, H, EP, 20060101, 20100224, A, L)
    B01D-15/00(B, I, M, 98, 20060101, 20100224, C)
    B01J-20/04(B, I, H, EP, 20060101, 20100224, A, L)
    B01J-20/04(B, I, M, 98, 20060101, 20100224, C)
Current IPC: A61K-39/09(B,I,H,EP,20060101,20100224,A,L)
    A61K-39/09(B, I, H, EP, 20100101, 20100224, C, L)
    A61K-47/48(B, I, H, EP, 20060101, 20100224, A, F)
    A61K-47/48(B, I, H, EP, 20100101, 20100224, C, F)
    A61P-31/00(B,I,H,EP,20100101,20100224,C,L)
    A61P-31/04(B, I, H, EP, 20060101, 20100224, A, L)
    B01D-15/00(B,I,H,EP,20060101,20100224,A,L)
    B01D-15/00(B, I, H, EP, 20100101, 20100224, C, L)
    B01J-20/04(B, I, H, EP, 20060101, 20100224, A, L)
    B01J-20/04(B, I, H, EP, 20100101, 20100224, C, L)
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Current ECLA ICO class: K61K-39:60P10
Original Abstract: This application relates to methods for the
    purification of saccharide antigen-carrier protein conjugates. In
    particular, the invention provides a method for purifying saccharide
    antigen- carrier protein conjugates from free carrier protein, such as
    CRM1 97, using hydroxyapatite. The invention further relates to methods
    of preparing vaccines, using this method.
United States
Publication No. US 20100239600 A1 (Update 201062 E)
Publication Date: 20100923
**CONJUGATE PURIFICATION**
Assignee: Novartis Vaccines and Diagnostics, Sienna, IT (NOVS)
  Bigio, Massimo, Siena, IT Residence: IT
  Averani, Giovanni, Siena, IT Residence: IT
  Norelli, Francesco, Siena, IT Residence: IT
  Berti, Francesco, Siena, IT Residence: IT
  Bellucci, Cinzia, Siena, IT Residence: IT
Inventor: Bigio, Massimo, Siena, IT Residence: IT
  Averani, Giovanni, Siena, IT Residence: IT
  Norelli, Francesco, Siena, IT Residence: IT
  Berti, Francesco, Siena, IT Residence: IT
Bellucci, Cinzia, Siena, IT Residence: IT
Agent: NOVARTIS VACCINES AND DIAGNOSTICS INC., INTELLECTUAL PROPERTY-
    X100B, P.O. BOX 8097, Emeryville, CA, US
Language: EN
Application: US 2010669464 A 20100609 (Local application)
  WO 2008IB2690 A 20080717 (PCT Application)
Priority: GB 200713880 A 20070717
Original IPC: A61K-39/385(B,I,H,US,20060101,20100923,A,F)
    A61K-39/385(B, I, M, 98, 20060101, 20100923, C)
    A61P-31/00(B, I, M, 98, 20060101, 20100923, C)
    A61P-31/12(B, I, H, US, 20060101, 20100923, A, L)
    A61P-37/00(B,I,H,US,20060101,20100923,A,L)
    A61P-37/00(B, I, M, 98, 20060101, 20100923, C)
    C07K-1/00(B, I, M, 98, 20060101, 20100923, C)
    C07K-1/14(B, I, H, US, 20060101, 20100923, A, L)
Current IPC: A61K-39/385(B,I,H,US,20060101,20100923,A,F)
    A61K-39/385(B, I, M, 98, 20060101, 20100923, C)
    A61P-31/00(B, I, M, 98, 20060101, 20100923, C)
    A61P-31/12(B, I, H, US, 20060101, 20100923, A, L)
    A61P-37/00(B, I, H, US, 20060101, 20100923, A, L)
    A61P-37/00(B, I, M, 98, 20060101, 20100923, C)
    C07K-1/00(B, I, M, 98, 20060101, 20100923, C)
    C07K-1/14(B, I, H, US, 20060101, 20100923, A, L)
Current ECLA class: A61K-39/09A A61K-47/48R2 A61K-47/48R2D A61K-47/48R2F
    A61K-47/48R2V C07K-1/16
Current ECLA ICO class: K61K-39:60P10
Current US Class (main): 424-193100
Current US Class (secondary): 530-351000 530-399000 530-413000
Original US Class (main): 424193.1
Original US Class (secondary): 530413 530351 530399
Original Abstract: This application relates to methods for the
    purification of saccharide antigen-carrier protein conjugates. In
    particular, the invention provides a method for purifying saccharide
    antigen- carrier protein conjugates from free carrier protein, such as
    CRM1 97, using hydroxyapatite. The invention further relates to methods
    of preparing vaccines, using this method.
Claim:
**1**. A method of purifying saccharide antigen-carrier protein
        conjugates from a mixture comprising free carrier protein and
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saccharide antigen-carrier protein conjugates, comprising contacting said mixture with hydroxyapatite such that the carrier protein binds to the hydroxyapatite while the conjugates do not

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bind; and collecting the free saccharide antigen-carrier protein
       conjugates.
Publication No. WO 2009010877 A2 (Update 200914 B)
Publication Date: 20090122
**CONJUGATE PURIFICATION
  PURIFICATION DE CONJUGUES**
Assignee: ~(only US)~ BIGIO, Massimo, Novartis Vaccines, Via Fiorentina 1,
    I-53100 Siena, IT Residence: IT Nationality: IT
  ~(only US)~ AVERANI, Giovanni, Novartis Vaccines, Via Fiorentina 1,
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  ~(only US)~ BERTI, Francesco, Novartis Vaccines, Via Fiorentina, 1,
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Inventor: AVERANI, Giovanni, Novartis Vaccines, Via Fiorentina 1, I-53100
    Siena, IT Residence: IT Nationality: IT
  BELLUCCI, Cinzia, Novartis Vaccines, Via Fiorentina, 1, I-53100 Siena, IT
    Residence: IT Nationality: IT
  BERTI, Francesco, Novartis Vaccines, Via Fiorentina, 1, I-53100 Siena, IT
    Residence: IT Nationality: IT
  BIGIO, Massimo, Novartis Vaccines, Via Fiorentina 1, I-53100 Siena, IT
    Residence: IT Nationality: IT
  NORELLI, Francesco, Novartis Vaccines, Via Fiorentina, 1, I-53100 Siena,
    IT Residence: IT Nationality: IT
Agent: MARSHALL, Cameron, John, Carpmaels Ransford, 43-45 Bloomsbury
    Square, London WC1A 2RA, GB
Language: EN (54 pages, 8 drawings)
Application: WO 2008IB2690 A 20080717 (Local application)
Priority: GB 200713880 A 20070717
Designated States: (National Original) AE AG AL AM AO AT AU AZ BA BB BG BH
    BR BW BY BZ CA CH CN CO CR CU CZ DE DK DM DO DZ EC EE EG ES FI GB GD GE
    GH GM GT HN HR HU ID IL IN IS JP KE KG KM KN KP KR KZ LA LC LK LR LS LT
    LU LY MA MD ME MG MK MN MW MX MY MZ NA NG NI NO NZ OM PG PH PL PT RO RS
    RU SC SD SE SG SK SL SM ST SV SY TJ TM TN TR TT TZ UA UG US UZ VC VN ZA
    ZM ZW
  (Regional Original) AT BE BG CH CY CZ DE DK EE ES FI FR GB GR HR HU IE IS
    IT LT LU LV MC MT NL NO PL PT RO SE SI SK TR OA BW GH GM KE LS MW MZ NA
    SD SL SZ TZ UG ZM ZW EA
Original IPC: C12N(99,20060101,S)
Current IPC: C12N(99,20060101,S)
Current ECLA class: A61K-39/09A A61K-47/48R2 A61K-47/48R2D A61K-47/48R2F
    A61K-47/48R2V C07K-1/16
Current ECLA ICO class: K61K-39:60P10
Original Abstract: This application relates to methods for the
    purification of saccharide antigen-carrier protein conjugates. In
    particular, the invention provides a method for purifying saccharide
    antigen- carrier protein conjugates from free carrier protein, such as
    CRM1 97, using hydroxyapatite. The invention further relates to methods
    of preparing vaccines, using this method.
   L'invention concerne des procedes de purification de conjugues antigene
    saccharidique-proteine de support. L'invention concerne en particulier
    un procede de purification des conjugues antigene
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saccharidique-proteine de support d'une proteine de support libre, par

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exemple CRM1 97, a l'aide d'hydroxyapatite. L'invention concerne
    egalement des methodes de preparation de vaccins a l'aide dudit
   procede.
Publication No. WO 2009010877 A3 (Update 200976 E)
Publication Date: 20091119
**CONJUGATE PURIFICATION**
Assignee: NOVARTIS AG; CH (NOVS)
Inventor: AVERANI G, IT
  BELLUCCI C, IT
  BERTI F, IT
 BIGIO M, IT
 NORELLI F, IT
Language: EN
Application: WO 2008IB2690 A 20080717 (Local application)
Priority: GB 200713880 A 20070717
Designated States: (National Original) AE AG AL AM AO AT AU AZ BA BB BG BH
    BR BW BY BZ CA CH CN CO CR CU CZ DE DK DM DO DZ EC EE EG ES FI GB GD GE
    GH GM GT HN HR HU ID IL IN IS JP KE KG KM KN KP KR KZ LA LC LK LR LS LT
    LU LY MA MD ME MG MK MN MW MX MY MZ NA NG NI NO NZ OM PG PH PL PT RO RS
    RU SC SD SE SG SK SL SM ST SV SY TJ TM TN TR TT TZ UA UG US UZ VC VN ZA
    ZM ZW
  (Regional Original) AT BE BG CH CY CZ DE DK EE ES FI FR GB GR HR HU IE IS
    IT LT LU LV MC MT NL NO PL PT RO SE SI SK TR OA BW GH GM KE LS MW MZ NA
    SD SL SZ TZ UG ZM ZW EA
Original IPC: A61K-39/09(B, I, H, EP, 20060101, A, L)
    A61K-39/09(B, I, M, 98, 20060101, C) A61K-47/48(B, I, H, EP, 20060101, A, F)
    A61K-47/48(B,I,M,98,20060101,C) A61P-31/00(B,I,M,98,20060101,C)
    A61P-31/04(B,I,H,EP,20060101,A,L) B01D-15/00(B,I,H,EP,20060101,A,L)
    B01D-15/00(B,I,M,98,20060101,C) B01J-20/04(B,I,H,EP,20060101,A,L)
    B01J-20/04(B, I, M, 98, 20060101, C)
Current IPC: A61K-39/09(B,I,H,EP,20060101,20091119,A,L)
   A61K-39/09(B, I, H, EP, 20090101, 20091119, C, L)
   A61K-47/48(B, I, H, EP, 20060101, 20091119, A, F)
    A61K-47/48(B, I, H, EP, 20090101, 20091119, C, F)
    A61P-31/00(B, I, H, EP, 20090101, 20091119, C, L)
    A61P-31/04(B, I, H, EP, 20060101, 20091119, A, L)
    B01D-15/00(B, I, H, EP, 20060101, 20091119, A, L)
    B01D-15/00(B, I, H, EP, 20090101, 20091119, C, L)
    B01J-20/04(B, I, H, EP, 20060101, 20091119, A, L)
    B01J-20/04(B, I, H, EP, 20090101, 20091119, C, L)
Current ECLA ICO class: K61K-39:60P10
Original Abstract: This application relates to methods for the
   purification of saccharide antigen-carrier protein conjugates. In
    particular, the invention provides a method for purifying saccharide
    antigen- carrier protein conjugates from free carrier protein, such as
    CRM1 97, using hydroxyapatite. The invention further relates to methods
    of preparing vaccines, using this method.
            (Item 8 from file: 351)
 10/7/8
DIALOG(R)File 351:Derwent WPI
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0018544858
WPI ACC NO: 2009-A43254/200902
XRAM Acc No: C2009-020226
New kit comprises conjugates of Haemophilus influenzae type B (Hib) and
****Neisseria**** ****meningitidis**** ****capsular**** saccharides, useful
for preparing a vaccine for raising an immune response against meningitis
Patent Assignee: NOVARTIS AG (NOVS); CONTORNI M (CONT-I); COSTANTINO P
  (COST-I)
Inventor: CONTORNI M; COSTANTINO P
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Patent Family (9 patents, 122 countries)
Patent
                              Application
Number
                                             Kind
               Kind
                     Date
                              Number
                                                  Date
                                                           Update
WO 2008149238
                A2 20081211 WO 2008IB2121
                                            A 20080604
                                                           200902
                                            A 20080604
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WO 2008149238
                A3 20090806 WO 2008IB2121
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AU 2008259423
                A1 20081211 AU 2008259423
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                    20100217 EP 2008789070
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MX 2009013112
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US 20100203137
                A1
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                                              P 20070604 201053
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                                              A 20100325
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JP 2010529103
                TeT
                    20100826
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                              JP 2010510910
                                               A 20080604
Priority Applications (no., kind, date): US 2007933235 P 20070604; US
  2010451831 A 20100325
Patent Details
                         Pg Dwg Filing Notes
Number
              Kind Lan
WO 2008149238
                          2.8
               A2 EN
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   GH GM GT HN HR HU ID IL IN IS JP KE KG KM KN KP KR KZ LA LC LK LR LS LT
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   BR BW BY BZ CA CH CN CO CR CU CZ DE DK DM DO DZ EC EE EG ES FI GB GD GE
   GH GM GT HN HR HU ID IL IN IS JP KE KG KM KN KP KR KZ LA LC LK LR LS LT
   LU LY MA MD ME MG MK MN MW MX MY MZ NA NG NI NO NZ OM PG PH PL PT RO RS
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AII 2008259423
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                                   Based on OPI patent
EP 2152302
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                                   PCT Application WO 2008IB2121
                                   Based on OPI patent WO 2008149238
Regional Designated States, Original: AT BE BG CH CY CZ DE DK EE ES FI FR
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                                   Based on OPI patent WO 2008149238
                                   PCT Application WO 2008IB2121
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                                   Based on OPI patent WO 2008149238
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                                   PCT Application WO 2008IB2121
                A1
                                   Based on OPI patent WO 2008149238
US 20100203137
                A1
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                                   Related to Provisional US 2007933235
                                   PCT Application WO 2008IB2121
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PCT Application WO 2008IB2121

Based on OPI patent WO 2008149238

JP 2010529103

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TAT

Alerting Abstract WO A2

NOVELTY - A kit comprising (i) an aqueous component, comprising a conjugate of a Hib ****capsular**** saccharide, and (ii) a lyophilized component, comprising a conjugate of a ~N. ****meningitidis**** ~ ****capsular**** saccharide, is new.

DESCRIPTION - INDEPENDENT CLAIMS are: (1) a method for preparing a combined vaccine by combining (i) an aqueous component, comprising a conjugate of a ~H. influenzae ~ type B (Hib) ****capsular**** saccharide, and (ii) a lyophilized component, comprising a conjugate of a ~N. ****meningitidis**** ~ ****capsular**** saccharide; (2) a combined vaccine comprising (i) a conjugate of a ~H. influenzae ~ type B ****capsular**** saccharide, and (ii) a conjugate of a ~N. ****meningitidis**** ~ ****capsular**** saccharide, prepared by combining an aqueous ~H. influenzae ~ conjugate and a lyophilized ~N. ****meningitidis**** ~ conjugate; (3) a vaccine comprising conjugates of ****capsular**** saccharides from two or more ~N. ****meningitidis**** ~ serogroups and from Hib, in an oil-in-water emulsion; (4) a kit, for preparing a vaccine, comprising (i) an oil-in-water emulsion component and (ii) a lyophilized component, comprising conjugated ****capsular**** saccharides from more than one serogroup of ~N. ****meningitidis**** ~; (5) a method for preparing a vaccine by combining (i) an oil-in-water emulsion component; and (ii) a lyophilized component comprising conjugates of ****capsular**** saccharides from more than one serogroup of ~N. ****meningitidis**** ~ ; (6) a vaccine comprising conjugates of ~N. ****meningitidis**** ~ ****capsular**** saccharides in an oil-in-water emulsion, prepared by combining an oil-in-water emulsion component and lyophilized conjugates of ****capsular**** saccharides from more than one serogroup of ~N. ****meningitidis**** ~ ; and (7) a method of raising an immune response in a patient by administering to the patient any of the vaccine above. ACTIVITY - Immunostimulant; Neuroprotective. No biological data given. MECHANISM OF ACTION - Vaccine.

MECHANISM OF ACTION - Vaccine.
USE - The kit is useful for preparing a vaccine. The vaccine is useful for raising an immune response against meningitis (all claimed).
ADVANTAGE - The present invention provides improved vaccine formulations for Hib and meningococcal conjugates.

Technology Focus

BIOTECHNOLOGY - Preferred Kit/Vaccine/Method: The aqueous component includes an adjuvant or is unadjuvanted. The ~H. influenzae ~ conjugate is adsorbed to aluminum phosphate. Administration of the ~H. influenzae ~ conjugate results in an anti-polyribosylribitol phosphate (PRP) antibody concentration in a patient of ^> 0.15mu g/ml. The concentration of ~H. influenzae ~ conjugate in the aqueous component is 0.5-50mu g/ml. The ~H. influenzae ~ saccharide is conjugated to a carrier protein selected from CRM197, tetanus toxoid, and the outer membrane complex of ~N. ****meningitidis**** ~ . The aqueous component comprises one or more of: a diphtheria toxoid, a tetanus toxoid, acellular pertussis antigen(s), inactivated poliovirus antigen(s), hepatitis B virus surface antigen, and/or pneumococcal saccharide. Administration of the ~N. ****meningitidis**** ~ conjugate(s) results in a bactericidal antibody response. The lyophilized component includes 2, 3, or 4 of meningococcal serogroups A, C, W135 and Y. The quantity of meningococcal ****capsular**** saccharide per serogroup is 1-20mu g. The ~N. ****meningitidis**** ~ saccharide(s) is/are conjugated to a carrier protein selected from CRM197, diphtheria toxoid and tetanus toxoid. The lyophilized component includes ****capsular**** includes a stabilizer. The lyophilized component does not include a Hib saccharide. The vaccine includes one or more buffers. The vaccine comprises one or more of: a diphtheria toxoid, a tetanus toxoid, acellular pertussis antigen(s), inactivated poliovirus antigen(s), hepatitis B virus surface antigen, and/or pneumococcal saccharide.

Title Terms/Index Terms/Additional Words: NEW; KIT; COMPRISE; CONJUGATE; HAEMOPHILUS; INFLUENZAE; TYPE; ****NEISSERIA****; ****MENINGITIDIS****; CAPSULE; USEFUL; PREPARATION; VACCINE; RAISE; IMMUNE; RESPOND; MENINGITIS

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Class Codes
International Classification (+ Attributes)
IPC + Level Value Position Status Version
 A61K-0039/095 A I F B 20060101
 A61K-0039/095 A I L B 20060101
 A61K-0039/102 A I F B 20060101
 A61K-0039/102 A I L B 20060101
 A61K-0039/116 A I L B 20060101
 A61K-0039/295 A I L B 20060101
 A61K-0039/39 A I L B 20060101
 A61K-0047/48 A I L B 20060101
 A61K-0009/107 A I F B 20060101
 A61K-0009/107 A I L B 20060101
 A61K-0009/19 A I L B 20060101
 A61P-0031/04 A I L B 20060101
A61K-0039/095 C I F B 20090101
  A61K-0039/095 C I
                       B 20060101
 A61K-0039/102 C I L B 20090101
A61K-0039/102 C I B 20060101
                        B 20060101
B 20060101
  A61K-0039/116 C I
 A61K-0039/295 C I L B 20100101
A61K-0039/39 C I B 20060101
 A61K-0047/48 C I L B 20090101
A61K-0009/107 C I F B 20100101
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 A61K-0009/19 C I L B 20090101
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 C12N S
ECLA: A61K-039/095, A61K-039/102, A61K-039/116, A61K-047/48R2V
ICO: K61K-039:545
US Classification, Current Main: 424-484000; Secondary: 424-201100
US Classification, Issued: 424484, 424201.1
JP Classification
 FI Term
                    Facet Rank Type
A61K-039/102
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A61K-039/095
                           B secondary
A61K-039/116
                          B secondary
A61K-039/39
                          B secondary
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A61P-031/04
A61K-039/095
A61K-039/102
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A61K-039/39
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F-Term View Point Additional
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4C085

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           GG01
File Segment: CPI
DWPI Class: B04: D16
Manual Codes (CPI/A-M): B04-B04C1; B04-C02F; B04-N03; B14-A01; B14-G01;
  B14-N16; B14-S11B1; D05-H07
Original Publication Data by Authority
Australia
Publication No. AU 2008259423 A1 (Update 201002 E)
Publication Date: 20081211
Assignee: NOVARTIS AG (NOVS)
Inventor: CONTORNI M
 COSTANTINO P
Language: EN
Application: AU 2008259423 A 20080604 (Local application)
Priority: US 2007933235 P 20070604
Related Publication: WO 2008149238 A (Based on OPI patent )
Original IPC: A61K-39/095(B,I,H,EP,20060101,20090806,A,F)
    A61K-39/102(B, I, H, EP, 20060101, 20090806, A, L)
    A61K-47/48(B, I, H, EP, 20060101, 20090806, A, L)
    A61K-9/107(B, I, H, EP, 20060101, 20090806, A, L)
    A61K-9/19 (B, I, H, EP, 20060101, 20090806, A, L)
Current IPC: A61K-39/095(B,I,H,EP,20060101,20090806,A,F)
    A61K-39/102(B, I, H, EP, 20060101, 20090806, A, L)
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    A61K-9/107(B, I, H, EP, 20060101, 20090806, A, L)
    A61K-9/19(B, I, H, EP, 20060101, 20090806, A, L)
Current ECLA class: A61K-39/095 A61K-39/102 A61K-39/116 A61K-47/48R2V
Current ECLA ICO class: K61K-39:545
Canada
Publication No. CA 2688268 A1 (Update 201020 E)
Publication Date: 20081211
Assignee: NOVARTIS AG; CH (NOVS)
Inventor: CONTORNI M, IT
  COSTANTINO P, IT
Language: EN
Application: CA 2688268 A 20080604 (Local application)
  WO 2008IB2121 A 20080604 (PCT Application)
  CA 2688268 A 20091125 (PCT national entry)
Priority: US 2007933235 P 20070604
Related Publication: WO 2008149238 A (Based on OPI patent )
Original IPC: A61K-39/095(B,I,H,EP,20060101,20090806,A,F)
    A61K-39/095(B, I, M, 98, 20060101, 20090806, C)
    A61K-39/102(B, I, H, EP, 20060101, 20090806, A, L)
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    A61K-47/48(B, I, H, EP, 20060101, 20090806, A, L)
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    A61K-9/107(B, I, H, EP, 20060101, 20090806, A, L)
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Current IPC: A61K-39/095(B,I,H,EP,20060101,20090806,A,F)
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    A61K-39/102(B, I, H, EP, 20060101, 20090806, A, L)
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Current ECLA class: A61K-39/095 A61K-39/102 A61K-39/116 A61K-47/48R2V
Current ECLA ICO class: K61K-39:545
China
Publication No. CN 101678094 A (Update 201024 E)
Publication Date: 20100324
**Formulation of meningitis vaccines**
Assignee: NOVARTIS AG; CH (NOVS)
Inventor: CONTORNI M, CH
 CONTORNI MARIO, CH
  COSTANTINO P. CH
  COSTANTINO PAOLO, CH
Language: ZH
Application: CN 200880018507 A 20080604 (Local application)
  WO 2008IB2121 A 20080604 (PCT Application)
Priority: US 2007933235 P 20070604
Related Publication: WO 2008149238 A (Based on OPI patent )
Original IPC: A61K-39/095(I.CN,20060101,A,F) A61K-39/095(I,M,98,20060101,C)
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    A61K-47/48(I,CN,20060101,A,L) A61K-47/48(I,M,98,20060101,C)
    A61K-9/107(I,CN,20060101,A,L) A61K-9/107(I,M,98,20060101,C)
    A61K-9/19(I,CN,20060101,A,L) A61K-9/19(I,M,98,20060101,C)
Current IPC: A61K-39/095(B,I,H,CN,20060101,20100325,A,F)
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    A61K-39/102(B, I, H, CN, 20100101, 20100325, C, L)
    A61K-47/48(B, I, H, CN, 20060101, 20100325, A, L)
    A61K-47/48(B, I, H, CN, 20100101, 20100325, C, L)
    A61K-9/107(B, I, H, CN, 20060101, 20100325, A, L)
    A61K-9/107(B, I, H, CN, 20100101, 20100325, C, L)
   A61K-9/19(B, I, H, CN, 20060101, 20100325, A, L)
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Current ECLA class: A61K-39/095 A61K-39/102 A61K-39/116 A61K-47/48R2V
Current ECLA ICO class: K61K-39:545
Original Abstract: A liquid Hib component is utilized to reconstitute a
    lyophilised meningococcal component, thereby producing a combined
    meningitis vaccine. A lyophilised meningococcal component can also be
    reconstituted with an oil-in-water emulsion.
Claim: [CLAIM 1] A kit, comprising: (i) an aqueous component, comprising a
    conjugate of a Haemophilus influenzae type B capsular saccharide; and
    (ii) a lyophilised component, comprising a conjugate of a Neisseria
    meningitidis capsular saccharide.
  [CLAIM 2] A method for preparing a combined vaccine, comprising the step
    of combining (i) an aqueous component, comprising a conjugate of a
    Haemophilus influenzae type B capsular saccharide, and (ii) a
    lyophilised component, comprising a conjugate of a Neisseria
    meningitidis capsular saccharide.
  [CLAIM 3] A combined vaccine, comprising: (i) a conjugate of a
    Haemophilus influenzae type B capsular saccharide; and (ii) a conjugate
    of a Neisseria meningitidis capsular saccharide, prepared by combining
    an aqueous H.influenzae conjugate and a lyophilised N. meningitidis
    conjugate.
  [CLAIM 4] The kit, method or vaccine according to any one of above
    claims, wherein the aqueous component includes an adjuvant.
  [CLAIM 5] The kit, method or vaccine according to any one of above
    claims, wherein the H.influenzae conjugate is adsorbed to aluminium
   phosphate.
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- [CLAIM 6] The kit, method or vaccine of any one of claims 1 to 3, wherein the agueous component is unadjuvanted.
- [CLAIM 7] A vaccine comprising conjugates of capsular saccharides from two or more Neisseria meningitidis serogroups and from Haemophilus influenzae type B, in an oil-in-water emulsion.
- [CLAIM 8] The kit, method or vaccine according to any one of above claims, wherein administration of the H.influenzae conjugate results in an anti-PRP antibody concentration in a patient of more than 0.15 micrograms/ml.
- [CLAIM 9] The kit, method or vaccine according to any one of above claims, wherein the concentration of H.influenzae conjugate in the aqueous component is in the range of 0.5 micrograms/ml to 50 micrograms/ml.
- [CLAIM 10] The kit, method or vaccine according to any one of above claims, wherein the H.influenzae saccharide is conjugated to a carrier protein selected from the group consisting of CRM1 97, tetanus toxoid, and the outer membrane complex of N. meningitidis.
- [CLAIM 11] The kit or method according to any one of above claims, wherein the aqueous component comprises one or more of: a diphtheria toxoid, a tetanus toxoid, acellular pertussis antigen(s), inactivated polioviras antigen(s), hepatitis B vims surface antigen, and/or pneumococcal saccharide.
- [CLAIM 12] A kit for preparing a vaccine, the kit comprising: (i) an oil-in-water emulsion component; and (ii) a lyophilised component, comprising conjugated capsular saccharides from more than one serogroup of Neisseria meningitidis.
- [CLAIM 13] A method for preparing a vaccine, comprising the step of combining: (i) an oil-in-water emulsion component; and (ii) a lyophilised component comprising conjugates of capsular saccharides from more than one seroqroup of Neisseria meningitidis.
- [CLAIM 14] A vaccine comprising conjugates of Neisseria meningitidis capsular saccharides in an oil-in-water emulsion, prepared by combining an oil-in-water emulsion component and lyophilised conjugates of capsular saccharides from more than one serogroup of N meningitidis.
- [CLAIM 15] The kit, method or vaccine according to any one of above claims, wherein administration of the N. meningitidis conjugate(s) results in a bactericidal antibody response.
- [CLAIM 16] The kit, method or vaccine according to any one of above claims, wherein the lyophilised component includes 2, 3, or 4 of meningococal serogroups A, C, W135 and Y.
- [CLAIM 17] The kit, method or vaccine of claim 16, wherein the lyophilised component includes capsular saccharides from each of meningococcal serogroups A, C, W135 and Y.
- [CLAIM 18] The kit, method or vaccine of claim 17, wherein the quantity of meningococcal capsular saccharide per serogroup is between 1 micrograms and 20 micrograms.
- [CLAIM 19] The kit, method or vaccine according to any one of above claims, wherein the N. meningitidis saccharide(s) is/are conjugated to a carrier protein selected from the group consisting of CRM1 97, diphtheria toxoid and tetanus toxoid.
- [CLAIM 20] The kit, method or vaccine of claim 16, wherein the lyophilised component includes capsular includes a stabiliser.
- [CLAIM 21] The kit, method or vaccine according to any one of above claims, wherein the lyophilised component includes an adjuvant.
- [CLAIM 22] The kit, method or vaccine of any one of claims 1 to 20, wherein the lyophilised component includes no adjuvant.
- [CLAIM 23] The kit, method or vaccine according to any one of above claims, wherein the lyophilised component does not include a Hib
- saccharide. [CLAIM 24] The vaccine according to any one of above claims, including one or more buffers.
- [CLAIM 25] The vaccine according to any one of above claims, wherein the

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vaccine comprises one or more of: a diphtheria toxoid, a tetanus
    toxoid, acellular pertussis antigen(s), inactivated poliovirus
    antigen(s), hepatitis B virus surface antigen, and/or pneumococcal
    saccharide.
  [CLAIM 26] A method of raising an immune response in a patient,
    comprising the step of administering to the patient a vaccine according
    to any one of above claims.
EPO
Publication No. EP 2152302 A2 (Update 201014 E)
Publication Date: 20100217
**FORMULIERUNG EINES IMPFSTOFFES GEGEN MENINGITIS
  FORMULATION OF MENINGITIS VACCINES
  FORMULATION DE VACCINS CONTRE LA MENINGITE**
Assignee: Novartis AG, Lichtstrasse 35, 4056 Basel, CH (NOVS)
Inventor: COSTANTINO, Paolo, Novartis Vaccines, Via Fiorentina 1, I-53100
   Siena, IT
  CONTORNI, Mario, Novartis Vaccines, Via Fiorentina 1, I-53100 Siena, IT
Agent: Marshall, Cameron John, Carpmaels Ransford, 43-45 Bloomsbury
    Square, London WC1A 2RA, GB
Language: EN
Application: EP 2008789070 A 20080604 (Local application)
  WO 2008IB2121 A 20080604 (PCT Application)
Priority: US 2007933235 P 20070604
Related Publication: WO 2008149238 A (Based on OPI patent )
Designated States: (Regional Original) AT BE BG CH CY CZ DE DK EE ES FI FR
    GB GR HR HU IE IS IT LI LT LU LV MC MT NL NO PL PT RO SE SI SK TR
Original IPC: A61K-39/095(B,I,H,EP,20060101,20091217,A,F)
    A61K-39/095(B, I, M, 98, 20060101, 20091217, C)
    A61K-39/102(B, I, H, EP, 20060101, 20091217, A, L)
    A61K-39/102(B, I, M, 98, 20060101, 20091217, C)
   A61K-47/48(B, I, H, EP, 20060101, 20091217, A, L)
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    A61K-9/107(B, I, H, EP, 20060101, 20091217, A, L)
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    A61K-9/19(B, I, M, 98, 20060101, 20091217, C)
Current IPC: A61K-39/095(B,I,H,EP,20060101,20091217,A,F)
    A61K-39/095(B, I, H, EP, 20100101, 20091217, C, F)
    A61K-39/102(B, I, H, EP, 20060101, 20091217, A, L)
    A61K-39/102(B, I, H, EP, 20100101, 20091217, C, L)
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    A61K-9/19(B, I, H, EP, 20060101, 20091217, A, L)
    A61K-9/19 (B, I, H, EP, 20100101, 20091217, C, L)
Current ECLA ICO class: K61K-39:545
Original Abstract: A liquid Hib component is used to reconstitute a
    lyophilised meningococcal component, thereby producing a combined
    meningitis vaccine. A lyophilised meningococcal component can also be
    reconstituted with an oil-in-water emulsion.
Japan
Publication No. JP 2010529103 W (Update 201056 E)
Publication Date: 20100826
Language: JA (33 pages)
Application: JP 2010510910 A 20080604 (Local application)
  WO 2008IB2121 A 20080604 (PCT Application)
Priority: US 2007933235 P 20070604
Related Publication: WO 2008149238 A (Based on OPI patent )
Original IPC: A61K-39/095(B,I,H,JP,20060101,20100730,A,L)
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    A61P-31/00(B, I, M, 98, 20060101, 20100730, C)
    A61P-31/04(B,I,H,JP,20060101,20100730,A,L)
Current IPC: A61K-39/095(B,I,H,JP,20060101,20100730,A,L)
    A61K-39/095(B, I, M, 98, 20060101, 20100730, C)
    A61K-39/102(B, I, H, JP, 20060101, 20100730, A, F)
    A61K-39/102(B, I, M, 98, 20060101, 20100730, C)
    A61K-39/116(B, I, H, JP, 20060101, 20100730, A, L)
    A61K-39/116 (B, I, M, 98, 20060101, 20100730, C)
    A61K-39/39(B, I, H, JP, 20060101, 20100730, A, L)
    A61K-39/39(B, I, M, 98, 20060101, 20100730, C)
    A61P-31/00(B, I, M, 98, 20060101, 20100730, C)
    A61P-31/04(B, I, H, JP, 20060101, 20100730, A, L)
Current ECLA class: A61K-39/095 A61K-39/102 A61K-39/116 A61K-47/48R2V
Current ECLA ICO class: K61K-39:545
Current JP FI-Terms: A61K-39/102 (main, A) A61K-39/095 (secondary, B)
    A61K-39/116 (secondary, B) A61K-39/39 (secondary, B) A61P-31/04
    (secondary, B) A61K-39/095 A61K-39/102 A61K-39/116 A61K-39/39
    A61P-31/04
Current JP F-Terms: 4C085 4C201 4C085AA03 4C085AA04 4C085AA38 4C085BA16
    4C085BA18 4C085EE03 4C085EE06 4C085FF01 4C085FF19 4C085GG01
Mexico
Publication No. MX 2009013112 A1 (Update 201028 E)
Publication Date: 20100131
Assignee: NOVARTIS AG (NOVS)
Inventor: CONTORNI M
 COSTANTINO P
Language: ES
Application: MX 200913112 A 20091202 (Local application)
  WO 2008IB2121 A 20080604 (PCT Application)
Priority: US 2007933235 P 20070604
Related Publication: WO 2008149238 A (Based on OPI patent )
Original IPC: A61K-39/095(I,MX,20060101,A,F) A61K-39/095(I,M,98,20060101,C)
    A61K-9/107(I,MX,20060101,A,L) A61K-9/107(I,M,98,20060101,C)
    A61K-9/19(I,MX,20060101,A,L) A61K-9/19(I,M,98,20060101,C)
Current IPC: A61K-39/095(B,I,H,MX,20100101,20060101,C,F)
    A61K-9/107(B, I, H, MX, 20100101, 20060101, C, L)
    A61K-9/19(B, I, H, MX, 20100101, 20060101, C, L)
Current ECLA class: A61K-39/095 A61K-39/102 A61K-39/116 A61K-47/48R2V
Current ECLA ICO class: K61K-39:545
United States
Publication No. US 20100203137 A1 (Update 201053 E)
Publication Date: 20100812
**FORMULATION OF MENINGITIS VACCINES**
Assignee: Contorni, Mario, Siena, IT Residence: IT (CONT-I)
  Costantino, Paolo, Colle Val d'Elsa, IT Residence: IT (COST-I)
Inventor: CONTORNI M, IT
 Costantino, Paolo, Colle Val d'Elsa, IT Residence: IT
Agent: NOVARTIS VACCINES AND DIAGNOSTICS INC., INTELLECTUAL PROPERTY-
    X100B, P.O. BOX 8097, Emeryville, CA, US
Language: EN
Application: US 2010451831 A 20100325 (Local application)
  WO 2008IB2121 A 20080604 (PCT Application)
 US 2007933235 P 20070604 (Related to Provisional)
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Original IPC: A61K-39/295(B,I,H,US,20060101,20100812,A,L)
    A61K-39/295(B, I, M, 98, 20060101, 20100812, C)
    A61K-9/107(B, I, H, US, 20060101, 20100812, A, F)
   A61K-9/107(B, I, M, 98, 20060101, 20100812, C)
Current IPC: A61K-39/295(B,I,H,US,20060101,20100812,A,L)
    A61K-39/295 (B, I, H, US, 20100101, 20100812, C, L)
    A61K-9/107(B, I, H, US, 20060101, 20100812, A, F)
    A61K-9/107(B, I, H, US, 20100101, 20100812, C, F)
Current ECLA class: A61K-39/095 A61K-39/102 A61K-39/116 A61K-47/48R2V
Current ECLA ICO class: K61K-39:545
Current US Class (main): 424-484000
Current US Class (secondary): 424-201100
Original US Class (main): 424484
Original US Class (secondary): 424201.1
Original Abstract: A liquid Hib component is used to reconstitute a
    lyophilised meningococcal component, thereby producing a combined
    meningitis vaccine. A lyophilised meningococcal component can also be
    reconstituted with an oil-in-water emulsion.
Claim:
**1**. A kit comprising: (i) an aqueous component, comprising a
       conjugate of a ~Haemophilus influenzae~ type B capsular saccharide;
        and (ii) a lyophilised component, comprising a conjugate of a
       ~Neisseria meningitidis~ capsular saccharide.
WIPO
Publication No. WO 2008149238 A2 (Update 200902 B)
Publication Date: 20081211
**FORMULATION OF MENINGITIS VACCINES
 FORMULATION DE VACCINS CONTRE LA MENINGITE**
Assignee: ~(except US)~ NOVARTIS AG, Lichtstrasse 35, CH-4056
Basel, CH Residence: CH Nationality: CH (NOVS)
  ~(only US)~ CONTORNI, Mario, Novartis Vaccines, Via Fiorentina 1, I-53100
    Siena, IT Residence: IT Nationality: IT
  ~(only US)~ COSTANTINO, Paolo, Novartis Vaccines, Via Fiorentina 1,
    I-53100 Siena, IT Residence: IT Nationality: IT
Inventor: COSTANTINO, Paolo, Novartis Vaccines, Via Fiorentina 1, I-53100
    Siena, IT Residence: IT Nationality: IT
  CONTORNI, Mario, Novartis Vaccines, Via Fiorentina 1, I-53100 Siena, IT
Agent: MARSHALL, Cameron, John, Carpmaels Ransford, 43-45 Bloomsbury
    Square, London WC1A 2RA, GB
Language: EN (28 pages, 0 drawings)
Application: WO 2008IB2121 A 20080604 (Local application)
Priority: US 2007933235 P 20070604
Designated States: (National Original) AE AG AL AM AO AT AU AZ BA BB BG BH
    BR BW BY BZ CA CH CN CO CR CU CZ DE DK DM DO DZ EC EE EG ES FI GB GD GE
    GH GM GT HN HR HU ID IL IN IS JP KE KG KM KN KP KR KZ LA LC LK LR LS LT
    LU LY MA MD ME MG MK MN MW MX MY MZ NA NG NI NO NZ OM PG PH PL PT RO RS
    RU SC SD SE SG SK SL SM SV SY TJ TM TN TR TT TZ UA UG US UZ VC VN ZA ZM
    ZW
  (Regional Original) AT BE BG BW CH CY CZ DE DK EA EE ES FI FR GB GH GM GR
    HR HU IE IS IT KE LS LT LU LV MC MT MW MZ NA NL NO OA PL PT RO SD SE SI
    SK SL SZ TR TZ UG ZM ZW
Original IPC: C12N(99,20060101,S)
Current IPC: C12N(99,20060101,S)
Current ECLA ICO class: K61K-39:545
Original Abstract: A liquid Hib component is used to reconstitute a
    lyophilised meningococcal component, thereby producing a combined
    meningitis vaccine. A lyophilised meningococcal component can also be
    reconstituted with an oil-in-water emulsion.
  Un composant liquide Hib est utilise pour reconstituer un composant a
   meningocoques lyophilise, afin de produire un vaccin combine contre la
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meningite. Le composant a meningocoques lyophilise peut aussi etre reconstitue a l'aide d'une emulsion huile dans l'eau.

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Publication No. WO 2008149238 A3 (Update 200952 E)
Publication Date: 20090806
**FORMULATION OF MENINGITIS VACCINES**
Assignee: NOVARTIS AG; CH (NOVS)
Inventor: CONTORNI M, IT
 COSTANTINO P, IT
Language: EN
Application: WO 2008IB2121 A 20080604 (Local application)
Priority: US 2007933235 P 20070604
Designated States: (National Original) AE AG AL AM AO AT AU AZ BA BB BG BH
    BR BW BY BZ CA CH CN CO CR CU CZ DE DK DM DO DZ EC EE EG ES FI GB GD GE
    GH GM GT HN HR HU ID IL IN IS JP KE KG KM KN KP KR KZ LA LC LK LR LS LT
    LU LY MA MD ME MG MK MN MW MX MY MZ NA NG NI NO NZ OM PG PH PL PT RO RS
    RU SC SD SE SG SK SL SM SV SY TJ TM TN TR TT TZ UA UG US UZ VC VN ZA ZM
  (Regional Original) AT BE BG CH CY CZ DE DK EE ES FI FR GB GR HR HU IE IS
    IT LT LU LV MC MT NL NO PL PT RO SE SI SK TR OA BW GH GM KE LS MW MZ NA
    SD SL SZ TZ UG ZM ZW EA
Original IPC: A61K-39/095(B,I,H,EP,20060101,A,F)
    A61K-39/095(B,I,M,98,20060101,C) A61K-39/102(B,I,H,EP,20060101,A,L)
    A61K-39/102(B,I,M,98,20060101,C) A61K-47/48(B,I,H,EP,20060101,A,L)
    A61K-47/48(B,I,M,98,20060101,C) A61K-9/107(B,I,H,EP,20060101,A,L)
    A61K-9/107(B,I,M,98,20060101,C) A61K-9/19(B,I,H,EP,20060101,A,L)
    A61K-9/19(B, I, M, 98, 20060101, C)
Current IPC: A61K-39/095(B,I,H,EP,20060101,20090806,A,F)
    A61K-39/095(B, I, H, EP, 20090101, 20090806, C, F)
    A61K-39/102(B, I, H, EP, 20060101, 20090806, A, L)
    A61K-39/102(B, I, H, EP, 20090101, 20090806, C, L)
    A61K-47/48(B, I, H, EP, 20060101, 20090806, A, L)
   A61K-47/48(B, I, H, EP, 20090101, 20090806, C, L)
   A61K-9/107(B, I, H, EP, 20060101, 20090806, A, L)
    A61K-9/107(B, I, H, EP, 20090101, 20090806, C, L)
    A61K-9/19(B, I, H, EP, 20060101, 20090806, A, L)
    A61K-9/19(B, I, H, EP, 20090101, 20090806, C, L)
Current ECLA ICO class: K61K-39:545
Original Abstract: A liquid Hib component is used to reconstitute a
    lyophilised meningococcal component, thereby producing a combined
    meningitis vaccine. A lyophilised meningococcal component can also be
   reconstituted with an oil-in-water emulsion.
 10/7/9
            (Item 9 from file: 351)
DIALOG(R)File 351:Derwent WPI
(c) 2011 Thomson Reuters. All rts. reserv.
0018017019 - Drawing available
WPI ACC NO: 2008-J37344/200854
XRAM Acc No: C2008-304663
Novel modified ****capsular**** saccharide, useful as medicament for
preventing or treating disease e.g. bacterial meningitis caused by
capsulate bacteria
Patent Assignee: NOVARTIS AG (NOVS); BARDOTTI A (BARD-I); BERTI F
  (BERT-I); COSTANTINO P (COST-I); PIANIGIANI A (PIAN-I)
Inventor: BARDOTTI A; BERTI F; COSTANTINO P; PIANIGIANI A
Patent Family (9 patents, 122 countries)
Patent
                               Application
Number
               Kind Date
                               Number
                                              Kind
                                                      Date
                                                              Update
WO 2008084411 A2 20080717 WO 2008IB1116 A 20080111 200854 B
WO 2008084411
               A3 20081224
                                                               200903 E
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AU 2008204259
               A1 20080717 AU 2008204259 A 20080111 200954 E
                                           A 20080111 200974 E
               A1 20080717
                            CA 2674228
CA 2674228
                                            A 20080111
                             WO 2008TB1116
                             CA 2674228
                                            A 20090630
                   20091118 EP 2008737594
EP 2118145
               A2
                                           A 20080111 200976
                             WO 2008IB1116 A 20080111
                            CN 200880002201 A 20080111 200978
CN 101583629
               A
                    20091118
                             WO 2008IB1116 A 20080111
MX 2009007415
               A1
                   20090930
                            WO 2008IB1116
                                            A 20080111 201007
                                            A 20090709
                             MX 20097415
JP 2010515718
                    20100513
                            WO 2008IB1116
                                            A 20080111 201032
                             JP 2009545256
                                            A 20080111
US 20100322958
              A1 20101223
                            WO 2008IB1116
                                            A 20080111 201101 E
                             US 2009448709
                                            A 20091106
Priority Applications (no., kind, date): GB 2007562 A 20070111
Patent Details
Number
             Kind Lan
                        Pg Dwg Filing Notes
WO 2008084411
               A2 EN
                         84
National Designated States, Original: AE AG AL AM AO AT AU AZ BA BB BG BH
  BR BW BY BZ CA CH CN CO CR CU CZ DE DK DM DO DZ EC EE EG ES FI GB GD GE
  GH GM GT HN HR HU ID IL IN IS JP KE KG KM KN KP KR KZ LA LC LK LR LS LT
  LU LY MA MD ME MG MK MN MW MX MY MZ NA NG NI NO NZ OM PG PH PL PT RO RS
  RU SC SD SE SG SK SL SM SV SY TJ TM TN TR TT TZ UA UG US UZ VC VN ZA ZM
Regional Designated States, Original: AT BE BG BW CH CY CZ DE DK EA EE ES
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  PL PT RO SD SE SI SK SL SZ TR TZ UG ZM ZW
WO 2008084411
              A3 EN
National Designated States, Original: AE AG AL AM AO AT AU AZ BA BB BG BH
  BR BW BY BZ CA CH CN CO CR CU CZ DE DK DM DO DZ EC EE EG ES FI GB GD GE
  GH GM GT HN HR HU ID IL IN IS JP KE KG KM KN KP KR KZ LA LC LK LR LS LT
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RU SC SD SE SG SK SL SM SV SY TJ TM TN TR TT TZ UA UG US UZ VC VN ZA ZM ZM Regional Designated States,Original: AT BE BG BW CH CY CZ DE DK EA EE ES FI FR GB GH GR HR HU IE IS IT KE LS LT LU LV MC MT MW MZ NA NL NO OA PL PT RO SD SE SI SK SL SZ TR TZ UG ZM ZW

LU LY MA MD ME MG MK MN MW MX MY MZ NA NG NI NO NZ OM PG PH PL PT RO RS

AU 2008204259 A1 EN Based on OPI patent WO 2008084411 CA 2674228 A1 EN PCT Application WO 20081B116 PCT national entry CA 2674228 Based on OPI patent WO 2008084411 PCT Application WO 20081B116 PCT Application WO 20081B116 WO 2008084411 PCT Application WO 20081B116 WO 2008084411 Based on OPI patent WO 2008084411

Regional Designated States,Original: AT BE BG CH CY CZ DE DK EE ES FI FR
GB GR HR HU IE IS IT LI LT LU LV MC MT NL MO PL PT RO SE SI SK TR
CN 101583629 A ZH PCT Application WO 20081B1116
Based on OPI patent WO 2008084411

MX 2009007415 A1 ES PCT Application W0 20091B11116

JF 2010515718 W JA 74 PT Application W0 20091B1116

Based on OPI patent W0 20091B1116

Based on OPI patent W0 20091B1116

Based on OPI patent W0 20091B1116

US 20100322958 A1 EN PCT Application W0 20091B1116

Alerting Abstract WO A2

NOVELTY - A modified ****capsular**** saccharide, comprising a blocking group at a hydroxyl group position on monosaccharide units of the corresponding native ****capsular**** saccharide, is new.

DESCRIPTION - A modified ****capsular**** saccharide comprising a blocking group at a hydroxyl group position on monosaccharide units of the corresponding native ****capsular**** saccharide, where the blocking group

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C(0), S(0) or SO 2;
X=
Y=
        NR 1 R 2 or R 3 :
       1-6C alkyl substituted with
R 1 =
       hydroxyl, sulfhydryl and amine;
 R 2 =
       H or 1-6C alkvl; and
R 3 = 1-6C \text{ alkyl.}
  INDEPENDENT CLAIMS are included for the following: (1) saccharide of
formula (I); (2) modifying a ****capsular**** saccharide, involves
providing ****capsular**** saccharide having hydroxyl group on a
monosaccharide unit, and converting the hydroxyl group into a blocking
group; (3) modifying ~N. ****meningitidis**** ~ serogroup A
****polvsaccharide****, involves (a) providing a native ~N.
****meningitidis**** ~ serogroup A ****polysaccharide****, depolymerizing
and sizing the ****polysaccharide**** to provide an oligosaccharide, and
converting at least one hydroxyl group of the oligosaccharide into a
blocking group, or (b) providing a native ~N. ****meningitidis**** ~
serogroup A ****polysaccharide****, converting hydroxyl group of
****polysaccharide**** into a blocking group, and depolymerizing and sizing
the resulting ****polysaccharide****; (4) preparing the modified
****capsular**** saccharide is a total synthesis process; (5) modified
****capsular**** saccharide obtained by the above method; (6)
saccharide-protein conjugate of a modified saccharide; (7) making a
saccharide-protein conjugate, involves (a) providing a modified
****capsular**** saccharide, and conjugating the modified ****capsular****
saccharide to a protein through the terminal anomeric hydroxyl group or the
amino group derived from a terminal anomeric hydroxyl group, or (b)
providing a modified ****capsular**** saccharide, converting the pairs of
vicinal hydroxyl groups into aldehyde groups by oxidative cleavage, and
linking the modified ****capsular**** saccharide to a protein by reductive
amination; (8) saccharide-protein conjugate of modified saccharide obtained
by above method; (9) molecule comprising a saccharide moiety of formula
(I); and (10) pharmaceutical composition comprising a modified saccharide
and/or saccharide-protein conjugate and/or molecule, and a carrier.
In formula (I), T=
                                       group of formulae (A) or (B), or In
                                      formula (I) contained in the
                                      molecule, T=group of formulae (C) or
                                      (D);
                                       1-100, preferably 15-25;
 n=
 7.=
                                       OH, OAc or blocking group;
 0=
                                       OH, OAc or blocking group;
                                       -NH 2 , -NHE, -NE 1 E 2 , W 2 , or
                                      -O-D:
 E, E 1 , E 2 =
                                       nitrogen protecting groups, which
                                      may be same or different;
                                       oxygen protecting group;
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-OH or blocking group;

is of formula: -O-X-Y (Ia), -O-R 1 (Ib), is new.

W, W 1 , W 2 =

Z,Q= OH or blocking groups of formula -O-X-Y' (IIa) or -O-R 4 (IIb); O, NH, NE, S or Se; Y '= NR 2 R 4 : R 2 =H or 1-6C alkyl; and R 4 =1-4C alkylene-CH(O) or -1-5C alkvlene-NH-.

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src="http://imagesrv.dialog.com/imanager/getimage?ref=Icdeedd006fda11dd8742000083613 46f&f=351&type=PNG" width="465" height="747"/> ACTIVITY - Antibacterial; Neuroprotective. No biological data given.

MECHANISM OF ACTION - Vaccine.

USE - The modified saccharide, conjugate or molecule is useful as a medicament for preventing or treating a disease caused by one or more capsulate bacteria. The disease is bacterial meningitis (all claimed). ADVANTAGE - The modified ****capsular**** saccharide has improved stability to hydrolysis.

Technology Focus

BIOTECHNOLOGY - Preparation (claimed): The modified ****capsular**** saccharide is produced by total synthesis process, by forming glycosidic linkages between two or more monosaccharide units. Preferred Components: The saccharide comprises blocking group of formula (Ia). The ratio of blocking groups in the saccharide is 90:10. At least 10% of the monosaccharide units of the saccharide have blocking groups. The monosaccharide units of the saccharide have blocking groups. The corresponding native ****capsular**** saccharide comprises monosaccharide units linked by phosphodiester bonds. The corresponding native ****capsular**** saccharide is ~****Neisseria**** meningitides ~ serogroup A saccharide. The blocking group is 4- and/or 3-positions of the corresponding ~N. ****meningitidis**** ~ serogroup A saccharide. The blocking group is 4-positions of the corresponding ~N.****meningitidis**** ~ serogroup A saccharide. The modified ****capsular**** saccharide is an oligosaccharide. The modified saccharide comprises a terminal anomeric hydroxyl group or an amino group derived from a terminal anomeric hydroxyl group. The monosaccharide unit of the modified ****capsular**** saccharide, where two vicinal hydroxyl groups of the corresponding native ****capsular*** saccharide do not comprise blocking groups. The monosaccharide units of the saccharide comprise blocking groups, where R 1 is substituted with two vicinal hydroxyl groups. At least 1% of the Q groups are blocking groups. The blocking group is -OC(0)NR 1 R 2 . Preferred Method: The converting step involves reacting the ****capsular**** saccharide with a bifunctional reagent in an organic solvent, and reacting the product of reacting step with an amino compound of formula: HNR 1 R 2 . The blocking group is -OC(0)R 3 and reacting the ****capsular**** saccharide with [R 3 C(0)] 2 O in the presence of an imidazole catalyst. The ****capsular**** saccharide is ****capsular**** oligosaccharide. The ****capsular**** oligosaccharide is obtained by depolymerizing and sizing the corresponding native ****capsular**** ****polysaccharide***. The ****capsular*** saccharide in the providing step is a native ****capsular*******polysaccharide**** and the method

further involves in which the product of converting step is sized, thus providing a modified ****capsular**** oligosaccharide. The vicinal hydroxyl groups present in the blocking groups are converted into aldehyde groups in converting step. The conditions for oxidative cleavage are proportion of the vicinal hydroxyl groups present in the blocking groups are converted into aldehyde groups in converting step. Preferred Conjugate: The protein is bacterial toxin or toxoid. The bacterial toxin or toxoid is diphtheria toxin or toxoid. The bacterial toxin or toxoid for the protein composition: The pharmaceutical composition further comprises saccharide antigen from one or more of serogroups C/MI35 and Y of *N.

****meningitidis***** o, the saccharide optionally being an oligosaccharide and optionally being conjugated to a carrier protein. The composition further comprises a vaccine adjuvant. The adjuvant is an aluminum phosphate. The composition is a vaccine against a disease caused by *N.

*****meningitidis*****.

Title Terms/Index Terms/Additional Words: NOVEL; MODIFIED; CAPSULE; SACCHARIDE; USEFUL; BEDICAMENT; PREVENT; TREAT; DISEASE; BACTERIA; MENINGITIS; CAUSE

Class Codes

International Classification (+ Attributes) IPC + Level Value Position Status Version A61K-0031/70 A I L B 20060101 A61K-0031/7024 A I L B 20060101 A61K-0031/715 A I L B 20060101 A61K-0038/16 A I L B 20660101 A61K-0039/02 A I L B 20660101 A61K-0039/095 A I L B 20060101 A61K-0039/095 A I F B 20060101 A61K-0039/385 A I L B 20060101 A61K-0039/39 A I L B 20060101 A61K-0047/48 A I L B 20060101 A61P-0031/04 A I L B 20060101 A61P-0037/04 A I L B 20060101 C07H-0011/00 A I L B 20060101 C07H-0011/04 A I F B 20060101 C07H-0011/04 A I L B 20060101 C07H-0013/02 A I L B 20060101 C07H-0013/12 A I L B 20060101 C07H-0015/04 A I L B 20060101 C07H-0005/06 A I L B 20060101 C07K-0014/00 A I L B 20060101 C07K-0014/195 A I L B 20060101 C07K-0014/34 A I L B 20060101 C08B-0037/00 A I F B 20060101 C08B-0037/00 A I L B 20060101 A61K-0031/70 C I B 20060101 A61K-0031/7024 C I B 2006010 A61K-0031/715 C I B 2006010 B 20060101 A61K-0031/7024 C I B 20060101 B 20060101 A61K-0038/16 C I B 20060101 A61K-0039/02 C I L B 20090101 A61K-0039/095 C I L B 20100101 A61K-0039/095 C I B 20060101 A61K-0039/385 C I B 20060101 AGIR-0039/385 C I B 20060101: AGIK-0039/39 C I L B 20090101 AGIR-0031/00 C I L B 20090101 AGIP-0031/00 C I B 20060101 AGIP-0031/00 C I B 20060101 COTH-0011/00 C I F B 20100101 C07H-0011/00 C I B 20060101

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C07H-0013/00 C I B 20060101
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C07H-0005/00 C I B 20060101
  C07K-0014/00 C I
                        B 20060101
  C07K-0014/195 C I
                         B 20060101
  C08B-0037/00 C I F B 20090101
  C08B-0037/00 C I L B 20100101
  C13K S
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ECLA: A61K-039/095, A61K-047/48R2V, C07H-013/00, C08B-037/00, C08B-037/00P
ICO: K61K-039:555A, K61K-039:60P10
US Classification, Current Main: 424-193100; Secondary: 424-184100,
424-250100, 514-001100, 514-023000, 514-054000, 530-404000, 530-405000,
530-406000, 530-408000, 530-409000, 530-411000, 536-018700, 536-053000,
536-054000, 536-117000, 536-118000, 536-119000, 536-120000
US Classification, Issued: 424193.1, 536118, 536119, 53653, 53654, 536120,
  536117, 53618.7, 530408, 530409, 530411, 530404, 530405, 530406, 51423,
  424250.1, 424184.1, 5141.1, 51454
JP Classification
  FI Term
                    Facet Rank Type
C07H-011/04
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A61K-039/095
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A61K-039/39
                           B secondary
A61P-031/04
                           B secondary
C08B-037/00
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A61K-039/39
A61P-031/04
C08B-037/00
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File Segment: CPI
DWPI Class: B03; B04
Manual Codes (CPI/A-M): B04-C02; B05-B02A3; B14-A01; B14-N16; B14-S11;
 N05-D
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Australia
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Assignee: NOVARTIS AG (NOVS)
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  COSTANTINO P
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Original IPC: A61K-39/02(B,I,H,EP,20060101,20081224,A,L)
    A61K-47/48 (B, I, H, EP, 20060101, 20081224, A, L)
    C08B-37/00(B, I, H, EP, 20060101, 20081224, A, F)
Current IPC: A61K-39/02(B,I,H,EP,20060101,20081224,A,L)
    A61K-39/02(B, I, H, EP, 20060101, 20081224, C, L)
    A61K-47/48(B, I, H, EP, 20060101, 20081224, A, L)
    A61K-47/48(B, I, H, EP, 20060101, 20081224, C, L)
    C08B-37/00(B, I, H, EP, 20060101, 20081224, A, F)
    C08B-37/00(B,I,H,EP,20060101,20081224,C,F)
Current ECLA class: A61K-39/095 A61K-47/48R2V C07H-13/00 C08B-37/00
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Current ECLA ICO class: K61K-39:555A K61K-39:60P10
Canada
Publication No. CA 2674228 A1 (Update 200974 E)
Publication Date: 20080717
Assignee: NOVARTIS AG; CH (NOVS)
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Language: EN
Application: CA 2674228 A 20080111 (Local application)
  WO 2008IB1116 A 20080111 (PCT Application)
  CA 2674228 A 20090630 (PCT national entry)
Priority: GB 2007562 A 20070111
Related Publication: WO 2008084411 A (Based on OPI patent )
Original IPC: A61K-39/02(B,I,H,EP,20060101,20081224,A,L)
    A61K-39/02(B,I,M,98,20060101,20081224,C)
    A61K-47/48(B, I, H, EP, 20060101, 20081224, A, L)
    A61K-47/48(B, I, M, 98, 20060101, 20081224, C)
    C08B-37/00(B, I, H, EP, 20060101, 20081224, A, F)
    C08B-37/00(B, I, M, 98, 20060101, 20081224, C)
Current IPC: A61K-39/02(B,I,H,EP,20060101,20081224,A,L)
    A61K-39/02(B,I,H,EP,20060101,20081224,C,L)
    A61K-47/48(B, I, H, EP, 20060101, 20081224, A, L)
    A61K-47/48(B, I, H, EP, 20060101, 20081224, C, L)
    C08B-37/00(B, I, H, EP, 20060101, 20081224, A, F)
    C08B-37/00(B, I, H, EP, 20060101, 20081224, C, F)
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Current ECLA class: A61K-39/095 A61K-47/48R2V C07H-13/00 C08B-37/00
    C08B-37/00P
Current ECLA ICO class: K61K-39:555A K61K-39:60P10
China
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**Modified saccharides**
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Language: ZH
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Related Publication: WO 2008084411 A (Based on OPI patent )
Original IPC: A61K-39/02(I,CN,20060101,A,L) A61K-39/02(I,M,98,20060101,C)
    A61K-47/48(I,CN,20060101,A,L) A61K-47/48(I,M,98,20060101,C)
    C08B-37/00(I,CN,20060101,A,F) C08B-37/00(I,M,98,20060101,C)
Current IPC: A61K-39/02(B,I,H,CN,20060101,20091118,A,L)
    A61K-39/02(B, I, H, CN, 20090101, 20091118, C, L)
    A61K-47/48 (B, I, H, CN, 20060101, 20091118, A, L)
    A61K-47/48(B, I, H, CN, 20090101, 20091118, C, L)
    C08B-37/00(B,I,H,CN,20060101,20091118,A,F)
    C08B-37/00(B,I,H,CN,20090101,20091118,C,F)
Current ECLA class: A61K-39/095 A61K-47/48R2V C07H-13/00 C08B-37/00
    C08B-37/00P
Current ECLA ICO class: K61K-39:555A K61K-39:60P10
Original Abstract: Modified capsular saccharides comprising a blocking
    group at a hydroxyl group position on at least one of the
    monosaccharide units of the corresponding native capsular saccharide,
    wherein the blocking group is of the formula (Ia) or (Ib), -OX-Y (Ia)
    or -O-R1 (Ib), wherein X is C(O), S(O) or SO2; Y is NR1R2 or R3; R1 is
    C1-6 alkyl substituted with 1, 2 or 3 groups independently selected
    from hydroxyl, sulphydryl and amine; R2 is H or C1-6 alkyl; and R3 is
    C1-6 alkyl; processes for modifying a capsular saccharide with the
    blocking groups; saccharide-protein conjugates comprising the modified
    capsular saccharide; processes for making the saccharide-protein
    conjugates, pharmaceutical compositions comprising the modified
    capsular saccharides and/or saccharide-protein conjugates; and methods
    and uses of the same.
Claim: [CLAIM 1] A modified capsular saccharide comprising a blocking group
    at a hydroxyl group position on at least one of the monosaccharide
    units of the corresponding native capsular saccharide, wherein the
   blocking group is of the formula (Ia) or (Ib), -O-X-Y (Ia) -O-R1 (Ib),
    wherein X is C(O), S(O) or SO2; Y is NR'R2 or R3; R1 is C1-6 alkyl
    substituted with 1, 2 or 3 groups independently selected from hydroxyl,
    sulphydryl and amine; R2 is H or C6 alkyl; R3 is C1-6 alkyl.
  [CLAIM 2] The modified capsular saccharide according to claim 1, wherein
    the blocking group is of formula (Ia).
  [CLAIM 3] The modified capsular saccharide according to claim 2, wherein
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[CLAIM 5] The modified capsular saccharide according to claim 4, wherein R2 is H. [CLAIM 6] The modified capsular saccharide according to claim 4 or claim 5, wherein R1 is substituted with 1, 2 or 3 hydroxyl groups.

[CLAIM 4] The modified capsular saccharide according to claim 2 or claim

X is C(0).

3, wherein Y is NR1R2.

- [CLAIM 7] The modified capsular saccharide according to any one of claims 4 to 6, wherein R1 is substituted with a single group, this substitution being at the distallend of the C1-6 alkyl chain.
- [CLAIM 8] The modified capsular saccharide according to any one of claims 4 to 6, wherein R1 is substituted with two vicinal groups.
- [CLAIM 9] The modified capsular saccharide according to any one of claims
- 4 to 8, wherein the saccharide comprises a) at least one blocking group, wherein Rl is substituted with a single group, this substitution being at the distal end of the C1-6 alkyl chain and b) at least one blocking group, wherein Rl is substituted with two vicinal groups.
- [CLAIM 10] The modified capsular saccharide according to claim 9, wherein the ratio of blocking groups, wherein R1 is substituted with a single group to blocking groups, wherein R1 is substituted with two vicinal groups is 90: 10.
- [CLAIM 11] The modified capsular saccharide according to claim 2 or claim 3, wherein Y is R3.
- [CLAIM 12] The modified capsular saccharide according to claim 11, wherein R3 is CH3.
- [CLAIM 13] The modified capsular saccharide according to any one of the preceding claims, wherein at least 10% of the monosaccharide units of the saccharide have blocking groups.
- [CLAIM 14] The modified capsular saccharide according to any one of the preceding claims, wherein all the monosaccharide units of the saccharide have blocking groups.
- [CLAIM 15] The modified capsular saccharide according to any one of the preceding claims, wherein the corresponding native capsular saccharide comprises monosaccharide units linked by phosphodiester bonds.
- [CLAIM 16] The modified capsular saccharide according to claim 15, wherein the corresponding native capsular saccharide is a Neisseria
- meningitidis serogroup A saccharide. [CLAIM 17] The modified capsular saccharide according to claim 16, wherein the blocking group is at any of the 4- and/or 3-positions of the corresponding Neisseria meningitidis serogroup A saccharide.
- [CLAIM 18] The modified capsular saccharide according to claim 17, wherein the blocking group is at any of the 4-positions of the corresponding Neisseria meningitidis serogroup A saccharide.
- [CLAIM 19] The modified capsular saccharide according to any one of the preceding claims, wherein the modified capsular saccharide is an oligosaccharide.
- [CLAIM 20] The modified capsular saccharide according to any one of the preceding claims, wherein the modified saccharide comprises a terminal anomeric hydroxyl group or an amino group derived from a terminal anomeric hydroxyl group.
- [CLAIM 21] The modified capsular saccharide according to any one of claims 1 to 19, wherein there is at least one monosaccharide unit of the modified capsular saccharide where two vicinal hydroxyl groups of the corresponding native capsular saccharide do not comprise blocking groups.
- [CLAIM 22] The modified capsular saccharide according to any one of claims 1 to 19, wherein at least one of the monosaccharide units of the saccharide comprise blocking groups, wherein R1 is substituted with two vicinal hydroxyl groups.
- [CLAIM 23] Å saccharide of the formula, FORMULA, wherein T is of the formula (A) or (B), FORMULA (A), FORMULA (B), n is an integer from 1 to 100; each Z group is independently selected from OH, OAc or a blocking group as defined in claims 1 to 8 or 11 to 12; and each Q group is independently selected from OH, OAc or a blocking group as defined in claims 1 to 8 or 11 to 12; V is selected from -NH2, -NHB, -NE1B2, W2, or -O-D, where: E, El and E2 are nitrogen protecting groups, which may be the same or different, and D is an oxygen protecting group; W is selected from -OH or a blocking group as defined in any one of claims 1 to 8 or 11 to 12; Wlis selected from -OH or a blocking group as defined in any one of claims 1

- in any one of claims 1 to 8 or 11 to 12; W2 is selected from -OH or a blocking group as defined in any one of claims 1 to 8 or 11 to 12; and wherein at least one of the Z groups and/or at least one of the Q groups are blocking groups as defined in claims 1 to 8 or 11 to 12.
- [CLAIM 24] The saccharide according to claim 23, wherein at least 10% of the Z groups are blocking groups.
- [CLAIM 25] The saccharide according to claim 23 or claim 24, wherein n is an integer from 15 to 25.
- [CLAIM 26] The saccharide according to any one of claims 23 to 25, wherein at least 1% of the O groups are blocking groups.
- [CLAIM 27] A process for modifying a capsular saccharide comprising the steps of: (a) providing a capsular saccharide having at least one hydroxyl group on a monosaccharide unit; and (b) converting said at least one hydroxyl group into a blocking group as defined in claims 1 to 8 or 11 to 12.
- [CLAIM 28] The process according to claim 27, wherein the blocking group is -OC(0)NRIR2 and step (b) comprises the steps of: (b1) reacting the capsular saccharide with a bifunctional reagent in an organic solvent; and (b2) reacting the product of step (b2) with an amino compound of formula (1), HNRIR2 (I), wherein R1 and R2 are as defined in any of claims 1 to 8.
- [CLAIM 29] The process according to claim 27, wherein the blocking group is -OC(0)R3 and step (b) comprises the step of (b1) reacting the capsular saccharide with [(R3C(0)]20 in the presence of an imidazole catalyst.
- [CLAIM 30] The process according to claim 28 or claim 29, wherein the capsular saccharide in step (a) is a capsular oligosaccharide.
- [CLAIM 31] The process according to claim 30, wherein the capsular oligosaccharide is obtainable by depolymerising and sizing the corresponding native capsular polysaccharide.
- [CLAIM 32] The process of claim 30, wherein the capsular saccharide in step (a) is a native capsular polysaccharide and the process further comprises a step (c) in which the product of step (b) is sized, thereby providing a modified capsular oligosaccharide.
- [CLAIM 33] A process for modifying a Neisseria meningitidis serogroup A polysaccharide comprising the steps of; (a) providing a native Neisseria meningitidis serogroup A polysaccharide; (b) depolymerising and sizing said polysaccharide to provide an oligosaccharide; and (c) converting at least one hydroxyl group of the oligosaccharide into a blocking group, in accordance with any one of claims 27 to 29.
- [CLAIM 34] A process for modifying a Neisseria meningitidis serogroup A polysaccharide comprising the steps of: (a) providing a native Neisseria meningitidis serogroup A polysaccharide; (b) converting at least one hydroxyl group of the polysaccharide into a blocking group, in accordance with any one of claims 27 to 29; and (c) depolymerising and sizing the resulting polysaccharide.
- [CLAIM 35] A process for preparing the modified capsular saccharide of claims 1 to 26 which is a total synthesis process comprising forming glycosidic linkages between two or more monosaccharide units.
- [CLAIM 36] A modified capsular saccharide obtainable or obtained by the process according to any one of claims 27 to 35.
- [CLAIM 37] A saccharide-protein conjugate of a modified saccharide according to any one of claims 1 to 26 or 36.
- [CLAIM 38] The conjugate according to claim 37, wherein the protein is a bacterial toxin or toxoid.
- [CLAIM 39] The conjugate according to claim 38, wherein the bacterial toxin or toxoid is diphtheria toxin or toxoid
- [CLAIM 40] The conjugate according to claim 39, wherein the bacterial toxin or toxoid is CRM197.
- [CLAIM 41] A process for making a saccharide-protein conjugate comprising the steps of: (c) providing a modified capsular saccharide according to claim 20; and (d) conjugating the modified capsular saccharide to a

- protein via the terminal anomeric hydroxyl group or the amino group derived from a terminal anomeric hydroxyl group.
- [CLAIM 42] A process for making a saccharide-protein conjugate comprising the steps of: (a) providing a modified capsular saccharide according to claim 21; (b) converting at least one of the pairs of vicinal hydroxyl groups into aldehyde groups by oxidative cleavage; and (c) linking the modified capsular saccharide to a protein by reductive amination.
- [CLAIM 43] A process for making a saccharide-protein conjugate comprising the steps of: (a) providing a modified capsular saccharide according to claim 22; (b) converting at least one of the pairs of vicinal hydroxyl groups into an aldehyde groups by oxidative cleavage; and (c) linking the modified capsular saccharide to a protein by reductive amination.
- [CLAIM 44] The process according to claim 43, wherein all of the vicinal hydroxyl groups present in the blocking groups are converted into aldehyde groups in step (b).
- [CLAIM 45] The process according to claim 43, wherein the conditions for oxidative cleavage are selected such that only a proportion of the vicinal hydroxyl groups present in the blocking groups are converted into aldehyde groups in step (b).
- [CLAIM 46] The process according to any one of claims 41 to 45, wherein the protein is as defined in any one of claims 38 to 40.
- [CLAIM 47] A saccharide-protein conjugate of a modified saccharide obtainable or obtained by the process according to any one of claims 41 to 46.
- CLAIM 48] A molecule comprising a saccharide moiety of formula, FORMULA, wherein T is of the formula (A) or (B), FORMULAE, n is an integer from 1 to 100, each Z group is independently selected from OH or a blocking group as defined in claims 1 o 8 or 11 to 12; and each Q group is independently selected from OH or a blocking group as defined in claims 1 o 8 or 11 to 12; W is selected from OH or a blocking group as defined in claims 1 to 8 or 11 to 12; L is O, NH, NE, S or Se, wherein the free covalent bond of L is joined to a protein carrier; and wherein the protein carrier; as defined in any one of claims 38 to 40, and wherein at least one of the Z groups and/or at least one of the Q groups are blocking groups as defined in claims 1 to 8 or 11 to 12.
- [CLAIM 49] A molecule comprising a saccharide of the formula, FORNULA, wherein T is of the formula (A) or (B), FORNULAE, n, Z, Q, W, W and V are as defined in claim 23, and at least one of the Z groups and/or at least one of the Q groups are of the formula -O-X-Y'(IIa), -O-R4 (IIb), wherein X is C(O), S(O) or SO2; Y'is NRZ4; R2 is H or C1-6alkyl; and R4 is -C1-4 alkylene-CH(O) or -C1-5 alkylene-NH-, wherein the -NH-group is part of a protein carrier; and wherein the protein carrier is a protein defined in any one of claims 38 to 40.
- [CLAÎM 50] A pharmaceutical composition comprising (a) a modified saccharide according to any one of claims 1 to 26 or 36 and/or a saccharide-protein conjugate according to any one of claims 37 to 40 or 47 and/or a molecule according to claim 48 or 49, and (b) a pharmaceutically acceptable carrier.
- [CLAIM 51] The composition according to claim 50, further comprising a saccharide antigen from one or more of serogroups C, W135 and Y of N. meningitidis, the saccharide optionally being an oligosaccharide and optionally being conjugated to a carrier protein.
- [CLAIM 52] The composition according to claim 50 or claim 51, further comprising a vaccine adjuvant.
- [CLAIM 53] The composition according to claim 52, wherein the adjuvant is an aluminium phosphate.
- [CLAIM 54] The composition according to any one of claims 50 to 53, which is a vaccine against a disease caused by N. meningitidis.
- [CLAIM 55] A method for raising an antibody response in a mammal,
- comprising administering the pharmaceutical composition according to any one of claims $50\ \text{to}54\ \text{to}$ the mammal.
- [CLAIM 56] The modified saccharide of any one according to claims 1 to 26

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or 36; the conjugate according to any one of claims 37 to 40 or 47; or
    the molecule according to claim 48 or 49 for use as a medicament.
  [CLAIM 57] The use of the modified polysaccharide according to any one of
    claims 1 to 26 or 36, or the conjugate according to any one of claims
    37 to 40 or 47, or the molecule according to claim 48 or 49, in the
    manufacture of a medicament for preventing or treating a disease caused
    by one or more capsulate bacteria.
  [CLAIM 58] The use according to claim 57, wherein the disease is
    bacterial meningitis.
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**MODIFIZIERTE SACCHARIDE
 MODIFIED SACCHARIDES
  SACCHARIDES MODIFIES**
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    GB GR HR HU IE IS ÎT LI LT LÛ LV MC MT NL NO PL PT RO SE SI SK TR
Original IPC: A61K-39/02(B,I,H,EP,20060101,20090812,A,L)
    A61K-39/02(B, I, M, 98, 20060101, 20090812, C)
    A61K-47/48(B, I, H, EP, 20060101, 20090812, A, L)
    A61K-47/48(B, I, M, 98, 20060101, 20090812, C)
    C08B-37/00(B, I, H, EP, 20060101, 20090812, A, F)
    C08B-37/00(B, I, M, 98, 20060101, 20090812, C)
Current IPC: A61K-39/02(B,I,H,EP,20060101,20090812,A,L)
    A61K-39/02(B, I, H, EP, 20090101, 20090812, C, L)
    A61K-47/48(B, I, H, EP, 20060101, 20090812, A, L)
    A61K-47/48(B, I, H, EP, 20090101, 20090812, C, L)
    C08B-37/00(B, I, H, EP, 20060101, 20090812, A, F)
    C08B-37/00(B, I, H, EP, 20090101, 20090812, C, F)
Current ECLA class: A61K-39/095 A61K-47/48R2V C07H-13/00 C08B-37/00
    C08B-37/00P
Current ECLA ICO class: K61K-39:555A K61K-39:60P10
Original Abstract: Modified capsular saccharides comprising a blocking
    group at a hydroxyl group position on at least one of the
    monosaccharide units of the corresponding native capsular saccharide,
    wherein the blocking group is of the formula (Ia) or (Ib): -OX-Y (Ia)
    or -0-R1 (Ib) wherein X is C(0), S(0) or SO2; Y is NR1R2or R3; R1 is
    C1-6 alkyl substituted with 1, 2 or 3 groups independently selected
    from hydroxyl, sulphydryl and amine; R2 is H or C1-6 alkyl; and R3 is
    C1-6 alkyl; processes for modifying a capsular saccharide with the
    blocking groups; saccharide-protein conjugates comprising the modified
    capsular saccharide; processes for making the saccharide-protein
    conjugates, pharmaceutical compositions comprising the modified
    capsular saccharides and/or saccharide-protein conjugates; and methods
    and uses of the same.
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Original IPC: A61K-39/095(B,I,H,JP,20060101,20100416,A,L)
    A61K-39/095(B,I,M,98,20060101,20100416,C)
    A61K-39/39(B, I, H, JP, 20060101, 20100416, A, L)
    A61K-39/39(B, I, M, 98, 20060101, 20100416, C)
    A61P-31/00(B, I, M, 98, 20060101, 20100416, C)
    A61P-31/04(B, I, H, JP, 20060101, 20100416, A, L)
    C07H-11/00(B, I, M, 98, 20060101, 20100416, C)
    C07H-11/04(B, I, H, JP, 20060101, 20100416, A, F)
    C08B-37/00(B, I, H, JP, 20060101, 20100416, A, L)
    C08B-37/00(B, I, M, 98, 20060101, 20100416, C)
Current IPC: A61K-39/095(B,I,H,JP,20060101,20100416,A,L)
    A61K-39/095(B, I, H, JP, 20100101, 20100416, C, L)
    A61K-39/39(B, I, H, JP, 20060101, 20100416, A, L)
    A61K-39/39(B,I,H,JP,20100101,20100416,C,L)
    A61P-31/00(B,I,H,JP,20100101,20100416,C,L)
    A61P-31/04(B,I,H,JP,20060101,20100416,A,L)
    C07H-11/00(B,I,H,JP,20100101,20100416,C,F)
    C07H-11/04(B, I, H, JP, 20060101, 20100416, A, F)
    C08B-37/00(B, I, H, JP, 20060101, 20100416, A, L)
    C08B-37/00(B, I, H, JP, 20100101, 20100416, C, L)
Current ECLA class: A61K-39/095 A61K-47/48R2V C07H-13/00 C08B-37/00
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Current ECLA ICO class: K61K-39:555A K61K-39:60P10
Current JP FI-Terms: C07H-11/04 (main, A, CSP) A61K-39/095 (secondary, B)
    A61K-39/39 (secondary, B) A61P-31/04 (secondary, B) C08B-37/00 P
    (secondary, B) C07H-11/04 (CSP) A61K-39/095 A61K-39/39 A61P-31/04
    C08B-37/00 P
Current JP F-Terms: 4C057 4C085 4C090 4C201 4C085AA03 4C090AA05 4C090AA09
    4C057AA17 4C057AA20 4C085AA38 4C085BA16 4C090BA94 4C090BA97 4C057BB02
    4C090BB64 4C090BB92 4C090BC25 4C090BD41 4C090CA35 4C057CC03 4C090DA23
    4C057DD03 4C085DD11 4C085DD59 4C085DD86 4C085EE06 4C085FF01 4C085FF13
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Inventor: BARDOTTI A
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  COSTANTINO P
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Original IPC: A61K-39/02(I,MX,20060101,A,L) A61K-39/02(I,M,98,20060101,C)
    A61K-47/48(I,MX,20060101,A,L) A61K-47/48(I,M,98,20060101,C)
    C08B-37/00(I,MX,20060101,A,F) C08B-37/00(I,M,98,20060101,C)
Current IPC: A61K-39/02(B,I,H,MX,20090101,20060101,C,L)
    A61K-47/48(B,I,H,MX,20090101,20060101,C,L)
    C08B-37/00(B, I, H, MX, 20090101, 20060101, C, F)
Current ECLA class: A61K-39/095 A61K-47/48R2V C07H-13/00 C08B-37/00
    C08B-37/00P
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**MODIFIED SACCHARIDES**
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Inventor: Bardotti, Angela, Siena, IT Residence: IT
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Agent: NOVARTIS VACCINES AND DIAGNOSTICS INC., INTELLECTUAL PROPERTY-
    X100B, P.O. BOX 8097, Emeryville, CA, US
Language: EN
Application: US 2009448709 A 20091106 (Local application)
  WO 2008IB1116 A 20080111 (PCT Application)
Priority: GB 2007562 A 20070111
Original IPC: A61K-31/70(B,I,H,US,20060101,20101223,A,L)
    A61K-31/70(B, I, M, 98, 20060101, 20101223, C)
    A61K-31/7024(B, I, H, US, 20060101, 20101223, A, L)
    A61K-31/7024(B, I, M, 98, 20060101, 20101223, C)
    A61K-31/715(B, I, H, US, 20060101, 20101223, A, L)
    A61K-31/715(B, I, M, 98, 20060101, 20101223, C)
    A61K-38/16(B, I, H, US, 20060101, 20101223, A, L)
    A61K-38/16(B, I, M, 98, 20060101, 20101223, C)
    A61K-39/095(B, I, H, US, 20060101, 20101223, A, F)
    A61K-39/095(B, I, M, 98, 20060101, 20101223, C)
    A61K-39/385(B, I, H, US, 20060101, 20101223, A, L)
    A61K-39/385 (B, I, M, 98, 20060101, 20101223, C)
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    A61P-31/04(B, I, H, US, 20060101, 20101223, A, L)
    A61P-37/00(B, I, M, 98, 20060101, 20101223, C)
    A61P-37/04(B, I, H, US, 20060101, 20101223, A, L)
    C07H-11/00(B, I, H, US, 20060101, 20101223, A, L)
    C07H-11/00(B, I, M, 98, 20060101, 20101223, C)
    C07H-11/04(B, I, H, US, 20060101, 20101223, A, L)
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    C07K-14/195(B, I, H, US, 20060101, 20101223, A, L)
    C07K-14/195(B, I, M, 98, 20060101, 20101223, C)
    C07K-14/34(B, I, H, US, 20060101, 20101223, A, L)
Current IPC: A61K-31/70(B,I,H,US,20060101,20101223,A,L)
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    A61K-31/7024(B, I, M, 98, 20060101, 20101223, C)
    A61K-31/715(B, I, H, US, 20060101, 20101223, A, L)
    A61K-31/715(B,I,M,98,20060101,20101223,C) A61K-38/
    16 (B, I, H, US, 20060101, 20101223, A, L)
    A61K-38/16(B, I, M, 98, 20060101, 20101223, C)
    A61K-39/095(B,I,H,US,20060101,20101223,A,F)
    A61K-39/095 (B, I, M, 98, 20060101, 20101223, C)
    A61K-39/385(B,I,H,US,20060101,20101223,A,L)
    A61K-39/385(B, I, M, 98, 20060101, 20101223, C)
    A61P-31/00(B, I, M, 98, 20060101, 20101223, C)
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A61P-31/04(B, I, H, US, 20060101, 20101223, A, L)
    A61P-37/00(B, I, M, 98, 20060101, 20101223, C)
    A61P-37/04(B,I,H,US,20060101,20101223,A,L)
    C07H-11/00(B, I, H, US, 20060101, 20101223, A, L)
   C07H-11/00(B,I,M,98,20060101,20101223,C)
    C07H-11/04(B, I, H, US, 20060101, 20101223, A, L)
   C07H-13/00(B, I, M, 98, 20060101, 20101223, C)
    C07H-13/02(B, I, H, US, 20060101, 20101223, A, L)
    C07H-13/12(B,I,H,US,20060101,20101223,A,L)
    C07H-15/00(B,I,M,98,20060101,20101223,C)
    C07H-15/04(B, I, H, US, 20060101, 20101223, A, L)
    C07H-5/00(B, I, M, 98, 20060101, 20101223, C)
    C07H-5/06(B, I, H, US, 20060101, 20101223, A, L)
    C07K-14/00(B, I, H, US, 20060101, 20101223, A, L)
    C07K-14/00(B, I, M, 98, 20060101, 20101223, C)
    C07K-14/195(B, I, H, US, 20060101, 20101223, A, L)
    C07K-14/195 (B, I, M, 98, 20060101, 20101223, C)
    C07K-14/34(B, I, H, US, 20060101, 20101223, A, L)
Current US Class (main): 424-193100
Current US Class (secondary): 424-184100 424-250100 514-001100 514-023000
    514-054000 530-404000 530-405000 530-406000 530-408000 530-409000
    530-411000 536-018700 536-053000 536-054000 536-117000 536-118000
    536-119000 536-120000
Original US Class (main): 424193.1
Original US Class (secondary): 536118 536119 53653 53654 536120 536117
    53618.7 530408 530409 530411 530404 530405 530406 51423 424250.1
    424184.1 5141.1 51454
Original Abstract: Modified capsular saccharides comprising a blocking
    group at a hydroxyl group position on at least one of the
    monosaccharide units of the corresponding native capsular saccharide,
    wherein the blocking group is of the formula (Ia) or (Ib): --OX--Y (Ia)
    or --O--R1 (Ib) wherein X is C(O), S(O) or SO2; Y is NR1R2 or R3; R1 is
    C1-6 alkyl substituted with 1, 2 or 3 groups independently selected
    from hydroxyl, sulphydryl and amine; R2 is H or C1-6 alkyl; and R3 is
    C1-6 alkyl; processes for modifying a capsular saccharide with the
    blocking groups; saccharide-protein conjugates comprising the modified
    capsular saccharide; processes for making the saccharide-protein
    conjugates, pharmaceutical compositions comprising the modified
    capsular saccharides and/or saccharide-protein conjugates; and methods
    and uses of the same.
Claim:
**1**. A modified capsular saccharide comprising a blocking group at
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- a hydroxyl group position on at least one of the monosaccharide units of the corresponding native capsular saccharide, wherein the blocking group is of the formula (Ia) or (Ib):
 - --O--X--Y (Ta)
 - * -- 0-- R1 (Ib)
 - * wherein
 - * X is C(0), S(0) or S02;
 - * Y is NR1R2 or R3:
 - * R1 is C1-6 alkyl substituted with 1, 2 or 3 groups independently selected from hydroxyl, sulphydryl and amine;
 - * R2 is H or C1-6 alkvl;
 - * R3 is C1-6 alkyl.

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WIPO
Publication No. WO 2008084411 A2 (Update 200854 B)
Publication Date: 20080717
**MODIFIED SACCHARIDES
  SACCHARIDES MODIFIES**
Assignee: ~(except US)~ NOVARTIS AG, Lightstrasse 35, CH-4056 Basel, CH
    Residence: CH Nationality: CH (NOVS)
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Language: EN (84 pages, 9 drawings)
Application: WO 2008IB1116 A 20080111 (Local application)
Priority: GB 2007562 A 20070111
Designated States: (National Original) AE AG AL AM AO AT AU AZ BA BB BG BH
    BR BW BY BZ CA CH CN CO CR CU CZ DE DK DM DO DZ EC EE EG ES FI GB GD GE
    GH GM GT HN HR HU ID IL IN IS JP KE KG KM KN KP KR KZ LA LC LK LR LS LT
    LU LY MA MD ME MG MK MN MW MX MY MZ NA NG NI NO NZ OM PG PH PL PT RO RS
    RU SC SD SE SG SK SL SM SV SY TJ TM TN TR TT TZ UA UG US UZ VC VN ZA ZM
  (Regional Original) AT BE BG BW CH CY CZ DE DK EA EE ES FI FR GB GH GM GR
    HR HU IE IS IT KE LS LT LU LV MC MT MW MZ NA NL NO OA PL PT RO SD SE SI
    SK SL SZ TR TZ UG ZM ZW
Original IPC: C13K(99,20060101,S)
Current IPC: C13K(99,20060101,S)
Current ECLA class: A61K-39/095 C07H-13/00 C08B-37/00 C08B-37/00P
Current ECLA ICO class: K61K-39:555A K61K-39:60P10
Original Abstract: Modified capsular saccharides comprising a blocking
    group at a hydroxyl group position on at least one of the
    monosaccharide units of the corresponding native capsular saccharide.
   wherein the blocking group is of the formula (Ia) or (Ib): -OX-Y (Ia)
    or -0-R1 (Ib) wherein X is C(O), S(O) or SO2; Y is NR1R2or R3; R1 is
    C1-6 alkyl substituted with 1, 2 or 3 groups independently selected
    from hydroxyl, sulphydryl and amine; R2 is H or C1-6 alkyl; and R3 is
    C1-6 alkyl; processes for modifying a capsular saccharide with the
    blocking groups; saccharide-protein conjugates comprising the modified
    capsular saccharide; processes for making the saccharide-protein
    conjugates, pharmaceutical compositions comprising the modified
    capsular saccharides and/or saccharide-protein conjugates; and methods
    and uses of the same.
  L'invention concerne des saccharides capsulaires modifies comprenant un
    groupe bloquant au niveau d'une position de groupe hydroxy sur au moins
    l'un des motifs monosaccharide du saccharide capsulaire natif
    correspondant, dans lesquels le groupe bloquant est de la formule (Ia)
    ou (Ib): -OX-Y (Ia) ou -O-R1 (Ib) dans lesquelles X represente C(O),
    S(O) ou SO2; Y represente NR1R2ou R3; R1 represente un groupe alkyle en
    Cl a C6 substitue par 1, 2 ou 3 groupes choisis independamment parmi le
    groupe hydroxy, sulfydryle et amine; R2 represente un atome H ou un
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groupe alkyle en C1 a C6; et R3 represente un groupe alkyle en C1 a C6; des procedes permettant de modifier un saccharide capsulaire avec les

groupes bloquants; des conjugues saccharide/proteine comprenant le saccharide capsulaire modifie; des procedes permettant de fabriquer les conjugues saccharide/proteine; des compositions pharmaceutiques comprenant des saccharides capsulaires modifies et/ou des conjugues saccharide/proteine, et des procedes dutilisation de ces compreses

saccharide/proteine: et des procedes d'utilisation de ces composes. Publication No. WO 2008084411 A3 (Update 200903 E) Publication Date: 20081224 Language: EN Designated States: (National Original) AE AG AL AM AO AT AU AZ BA BB BG BH BR BW BY BZ CA CH CN CO CR CU CZ DE DK DM DO DZ EC EE EG ES FI GB GD GE GH GM GT HN HR HU ID IL IN IS JP KE KG KM KN KP KR KZ LA LC LK LR LS LT LU LY MA MD ME MG MK MN MW MX MY MZ NA NG NI NO NZ OM PG PH PL PT RO RS RU SC SD SE SG SK SL SM SV SY TJ TM TN TR TT TZ UA UG US UZ VC VN ZA ZM 2W(Regional Original) AT BE BG BW CH CY CZ DE DK EA EE ES FI FR GB GH GM GR HR HU IE IS IT KE LS LT LU LV MC MT MW MZ NA NL NO OA PL PT RO SD SE SI SK SL SZ TR TZ UG ZM ZW Original IPC: A61K-39/02(B, I, H, EP, 20060101, A, L) A61K-39/02(B,I,M,98,20060101,C) A61K-47/48(B,I,H,EP,20060101,A,L) A61K-47/48(B,I,M,98,20060101,C) C08B-37/00(B,I,H,EP,20060101,A,F) C08B-37/00(B,I,M,98,20060101,C) Current IPC: A61K-39/02(B,I,H,EP,20060101,20081224,A,L) A61K-39/02(B,I,H,EP,20060101,20081224,C,L) A61K-47/48(B, I, H, EP, 20060101, 20081224, A, L) A61K-47/48(B,I,H,EP,20060101,20081224,C,L) C08B-37/00(B,I,H,EP,20060101,20081224,A,F) C08B-37/00(B,I,H,EP,20060101,20081224,C,F) Current ECLA class: A61K-39/095 C07H-13/00 C08B-37/00 C08B-37/00P Current ECLA ICO class: K61K-39:555A K61K-39:60P10 10/7/10 (Item 10 from file: 351) DIALOG(R)File 351:Derwent WPI (c) 2011 Thomson Reuters. All rts. reserv. 0017833268 WPI ACC NO: 2008-G53727/200841 XRAM Acc No: C2008-207379 XRPX Acc No: N2008-513921 Analyzing degree of unconjugation of a sample by contacting the sample with a basic reagent to precipitate conjugated saccharide component from sample and obtain supernatant comprising the unconjugated component Patent Assignee: NOVARTIS VACCINES & DIAGNOSTICS INC (NOVS); NOVARTIS VACCINES&DIAGNOSTICS SRL (NOVS) Inventor: BERTI F; COSTANTINO P; GALLETTI B; PARENTE P Patent Family (4 patents, 119 countries) Patent Application Number Kind Kind Date Date Number Update WO 2007122512 A2 20071101 WO 2007IB1855 A 20070321 200841 WO 2007122512 A3 20080124 200841 E EP 2005165 A2 20081224 EP 2007734937 A 20070321 200903 E A 20070321 WO 2007IB1855 A 20070321 200946 E US 20090176311 A1 20090709 WO 2007IB1855 A 20090218 US 2009293130

Priority Applications (no., kind, date): GB 20065757 A 20060322

Patent Details Number Kind Lan Pg Dwg Filing Notes WO 2007122512 A2 EN 33 3 National Designated States,Original: AE AG AL AM AT AU AZ BA BB BG BH BR BW BY BZ CA CH CN CO CR CU CZ DE DK DM DZ EC EE EG ES FI GB GD GE GH GM GT HN HR HU ID IL IN IS JP KE KG KM KN KP KR KZ LA LC LK LR LS LT LU LY MA MD MG MK MM MM MX MY MZ NA NG NI NO NZ OM PG PH PL PT RO RS RU SC S SE SG SK SL SM SV SY TJ TM TN TR TT TZ UA UG US UZ VC VN ZA ZM ZW REGional Designated States, Original: AT BE BG BW CH CY CZ DE DK EA EE ES FI FR GB GH GM GR HU IE IS IT KE LS LT LU LV MC MT MW MZ NA NL OA PL PT RO SD SE SI SK SL SZ TR TZ UG ZM ZW

National Designated States, Original: AE AG AL AM AT AU AZ BA BB BG BH BR BW BY BZ CA CH CN CO CR CU CZ DE DK DM DZ EC EE EG ES FI GB GD GE GH GM GT HN HR HU ID IL IN IS JP KE KG KM KN KP KR KZ LA LC LK LR LS LT LU LY MA MD MG MK MM MW MX MY MZ MA NG NI NO NZ OM PG PH PL PT RO RS RU SC SD SE SG SK KL SM SV SY TJ TM NI TR TT TZ UD UG US UZ VC VN ZA ZM ZW

Regional Designated States, Original: AT BE BG BW CH CY CZ DE DK EA EE ES FI FR GB GH GK HU IE IS IT KE LS LT LU LV MC MT MW MZ NA NL OA PL PT RO SD SE SI SK SL SZ TR TZ UG ZM ZW EP 2005165 A2 EN PCT Application WO 2007IB1855

Based on OPI patent WO 2007122512
Regional Designated States,Original: AT BE BG CH CY CZ DE DK EE ES FI FR
GB GR HU IE IS IT LI LT LU LV MC MT NL PL PT RO SE SI SK TR
US 20090176311 AI EN PCT Application WO 20071B1855

Alerting Abstract WO A2

NOVELTY — Analyzing a sample's degree of unconjugation comprises (a) contacting the sample with a basic reagent under basic conditions to selectively precipitate the conjugated saccharide component from the sample and to obtain a supernatant comprising the separated unconjugated component, and (b) analyzing the supernatant's content to give the unconjugated content of the sample.

DESCRIPTION - INDEPENDENT CLAIMS are:

- a method of preparing a sample for analysis of its degree of unconjugation;
- 2.a method of separating a conjugated saccharide component in a sample from an unconjugated component in the sample;
- 3.a supernatant obtained by the method above;
- 4.a precipitate obtained by the method above;
- 5.a method of releasing a vaccine for use by physicians;
- 6.a method for preparing a vaccine composition; and
- 7.a method for packaging a vaccine.

USE - The basic reagent under basic conditions used in the methods above is useful for selectively precipitating a conjugated saccharide component in a sample from an unconjugated component in the sample, thus separating the conjugated saccharide component from the unconjugated component (claimed). The methods are useful for analyzing a sample's degree of unconjugation, preparing a sample for analyzing of its degree of unconjugation, and separating a conjugated saccharide component in a sample from an unconjugated component in the sample. The methods can be used for the analysis and quality control of conjugate vaccines and for monitoring the stability of a vaccine in storage.

ADVANTAGE — The invention overcomes the deficiencies in the prior art and provides a rapid and quantitative technique for separation of unconjugated and conjugated components. It is applicable to a range of conjugate vaccines, including vaccines comprising bacterial ****capsular**** saccharides containing a sialic acid residue and particularly vaccines

comprising ~Streptococcus agalactiae ~ ****capsular**** saccharide.

Technology Focus

BIOTECHNOLOGY - Preferred Method: The improved method of analyzing the degree of unconjugation of a sample comprises contacting the sample with a basic reagent under basic conditions to selectively precipitate the conjugated saccharide component from the sample. Preparing a sample for analysis of its degree of unconjugation comprises contacting the sample with a basic reagent under basic conditions to selectively precipitate the conjugated saccharide component from the sample and to obtain a supernatant comprising separated unconjugated component. The unconjugated component is an unconjugated saccharide component or an unconjugated carrier component. It also comprises measuring the sample's total saccharide content or measuring the sample's total carrier content. In the method, the basic reagent comprises a lyotropic salt, where the lyotropic salt is a sulfate, hydrogen phosphate, acetate, citrate, tartrate of ammonium, potassium, sodium, or lithium. Preferably, the lyotropic salt is a sulfate or hydrogen phosphate of ammonium or potassium. Specifically, the lyotropic salt is K 2 HPO 4 . In the method, the basic conditions are from pH 8-12. Preferably, the basic reagent comprises K 2 HPO 4 and the basic conditions are from pH 9.5-9.9. Preferably, the conjugated saccharide is a saccharide antigen conjugated to a carrier protein. The sample is a vaccine, where the vaccine is a glycoconjugate vaccine. The glycoconjugate vaccine comprises a conjugate comprising a saccharide containing a sialic acid residue. It also comprises a conjugate comprising a bacterial ****capsular**** saccharide from ~Streptococcus agalactiae ~ . Preferably, the bacterial ****capsular*** saccharide is from ~S. agalactiae ~ serogroup Ia, Ib, II, III, or V. Separating a conjugated saccharide component in a sample from an unconjugated component in the sample comprises contacting the sample with a basic reagent under basic conditions to selectively precipitate the conjugated saccharide component from the sample. The improvement comprises contacting the sample with a basic reagent under basic conditions to selectively precipitate the conjugated saccharide component from the sample. Releasing a vaccine for use by physicians comprises:

- 1.manufacturing a vaccine comprising a conjugated saccharide;
- analyzing the vaccine's degree of unconjugation by the method above;
- 3.if the results from (b) indicate a degree of unconjugation acceptable for clinical use, releasing the vaccine for use by physicians.

Preparing a vaccine composition comprises analyzing the vaccine's degree of unconjugation by the method above including pH measurement, followed by adjusting the pH of the composition to a desired value, e.g. 6-8, preferably 7. Packaging a vaccine comprises:

- 1.manufacturing a bulk vaccine containing a conjugated saccharide;
- 2.analyzing the degree of unconjugation of the bulk vaccine by the method above;
- 3.optionally, analyzing the bulk vaccine for pH and/or other properties;
- 4.if the results from (b) and (c) indicate that the bulk vaccine is acceptable for clinical use, preparing and packaging the vaccine for human use from the bulk.

REAGENT; PRECIPITATION; CONJUGATE; SACCHARIDE; COMPONENT; OBTAIN; SUPERNATANT; COMPRISE; UNCONJUGATED

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Class Codes
International Classification (+ Attributes)
IPC + Level Value Position Status Version
 A61K-0039/00 A I L B 20060101
 C07H-0001/00 A I L B 20060101
 G01N-0033/15 A N L
                           20060101
 G01N-0033/50 A I F
                           20060101
 G01N-0033/50 A I F B 20060101
 A61K-0039/00 C I L B 20060101
 A61K-0039/00 C I
                        B 20060101
 C07H-0001/00 C I
                       B 20060101
 G01N-0033/15 C N
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 G01N-0033/50 C I
                           20060101
 G01N-0033/50 C I F B 20060101
 G01N-0033/50 C I
                        B 20060101
ICO: K61K-039:60P10
US Classification, Current Main: 436-094000; Secondary: 536-127000
US Classification, Issued: 43694, 536127
File Segment: CPI: EPI
DWPI Class: B04; D16; S03
Manual Codes (EPI/S-X): S03-E13D
Manual Codes (CPI/A-M): B04-B04C1; B04-C02F; B04-N06; B07-A02B; B11-C06;
 B11-C08D; B11-C08E; B12-K04A; B12-K04E3; B14-S11; D05-H07; D05-H09;
 D05-H13
Original Publication Data by Authority
EPO
Publication No. EP 2005165 A2 (Update 200903 E)
Publication Date: 20081224
**TRENNUNG KONJUGIERTER UND NICHTKONJUGIERTER KOMPONENTEN
 SEPARATION OF CONJUGATED AND UNCONJUGATED COMPONENTS
 SEPARATION DE COMPOSES CONJUGUES ET NON CONJUGUES**
Assignee: Novartis Vaccines and Diagnostics S.r.l., Via Fiorentina 1, 53100
   Siena (SI), IT
Inventor: BERTI, Francesco, Novartis Vaccines and Diagnostics S.r.l., Via
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Agent: Marshall, Cameron John, Carpmaels Ransford, 43-45 Bloomsbury
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Language: EN
Application: EP 2007734937 A 20070321 (Local application)
 WO 2007IB1855 A 20070321 (PCT Application)
Priority: GB 20065757 A 20060322
Related Publication: WO 2007122512 A (Based on OPI patent )
Designated States: (Regional Original) AT BE BG CH CY CZ DE DK EE ES FI FR
   GB GR HU IE IS IT LI LT LU LV MC MT NL PL PT RO SE SI SK TR
Original IPC: A61K-39/00(B,I,H,EP,20060101,20081029,A,L)
   A61K-39/00(B, I, M, 98, 20060101, 20081029, C)
   G01N-33/50(B, I, H, EP, 20060101, 20081029, A, F)
   G01N-33/50(B,I,M,98,20060101,20081029,C)
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Current IPC: A61K-39/00(B,I,H,EP,20060101,20081029,A,L)
    A61K-39/00(B, I, M, 98, 20060101, 20081029, C)
    G01N-33/50(B, I, H, EP, 20060101, 20081029, A, F)
    G01N-33/50(B,I,M,98,20060101,20081029,C)
Current ECLA ICO class: K61K-39:60P10
Original Abstract: The invention is based on the use of a basic reagent
    under basic conditions to separate conjugated saccharide from
    unconjugated components in a sample, ~e.g.~ a vaccine, by precipitation
    of the conjugated saccharide. The invention allows rapid and
    quantitative separation of conjugated and conjugated components, which
    may be exploited in analytical methods for quantifying unconjugated
    saccharide or carrier. Therefore, the separation of conjugated and
    unconjugated components using the invention may be advantageously
    combined with a quantitative saccharide or carrier analysis to provide
    improved quality control for conjugate vaccines.
United States
Publication No. US 20090176311 A1 (Update 200946 E)
Publication Date: 20090709
**SEPARATION OF CONJUGATED AND UNCONJUGATED COMPONENTS**
Assignee: NOAVARTIS VACCINES AND DIAGNOSTICS SRL, Siena, IT (NOVS)
  Berti, Francesco, Siena, IT Residence: IT
  Galletti, Bruno, Siena, IT Residence: IT
  Parente, Pierino, Siena, IT Residence: IT
  Costantino, Paolo, Siena, IT Residence: IT
Inventor: Berti, Francesco, Siena, IT Residence: IT
  Costantino, Paolo, Siena, IT Residence: IT
  Galletti, Bruno, Siena, IT Residence: IT
  Parente, Pierino, Siena, IT Residence: IT
Agent: NOVARTIS VACCINES AND DIAGNOSTICS INC., INTELLECTUAL PROPERTY R338,
    P.O. BOX 8097, Emeryville, CA, US
Language: EN
Application: US 2009293130 A 20090218 (Local application)
  WO 2007IB1855 A 20070321 (PCT Application)
Priority: GB 20065757 A 20060322
Original IPC: C07H-1/00(B,I,H,US,20060101,20090709,A,L)
    C07H-1/00(B, I, M, 98, 20060101, 20090709, C)
    G01N-33/50(B, I, H, US, 20060101, 20090709, A, F)
    G01N-33/50(B, I, M, 98, 20060101, 20090709, C)
Current IPC: C07H-1/00(B,I,H,US,20060101,20090709,A,L)
    C07H-1/00(B, I, M, 98, 20060101, 20090709, C)
    G01N-33/50(B, I, H, US, 20060101, 20090709, A, F)
    G01N-33/50(B, I, M, 98, 20060101, 20090709, C)
Current ECLA ICO class: K61K-39:60P10
Current US Class (main): 436-094000
Current US Class (secondary): 536-127000
Original US Class (main): 43694
Original US Class (secondary): 536127
Original Abstract: The invention is based on the use of a basic reagent
    under basic conditions to separate conjugated saccharide from
    unconjugated components in a sample, e.g. a vaccine, by precipitation
    of the conjugated saccharide. The invention allows rapid and
    quantitative separation of conjugated and conjugated components, which
    may be exploited in analytical methods for quantifying unconjugated
    saccharide or carrier. Therefore, the separation of conjugated and
    unconjugated components using the invention may be advantageously
    combined with a quantitative saccharide or carrier analysis to provide
    improved quality control for conjugate vaccines.
Claim:
**1**. A method of analysing a sample's degree of unconjugation,
        comprising the steps of (i) contacting the sample with a basic
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reagent under basic conditions to selectively precipitate the conjugated saccharide component from the sample and thereby to obtain a supernatant comprising the separated unconjugated component and (ii) analysing the supernatant's content to give the

WIPO

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unconjugated content of the sample.
Publication No. WO 2007122512 A2 (Update 200841 B)
Publication Date: 20071101
**SEPARATION OF CONJUGATED AND UNCONJUGATED COMPONENTS
  SEPARATION DE COMPOSES CONJUGUES ET NON CONJUGUES**
Assignee: ~(except US)~ NOVARTIS VACCINES AND DIAGNOSTICS SRL, Via
    Fiorentina 1, I-53100 Siena, IT Residence: IT Nationality: IT (NOVS)
  ~(only US)~ BERTI, Francesco, c/o Novartis Vaccines and Diagnostics
    S.r.l., Via Fiorentina, 1, I-53100 Siena, IT Residence: IT Nationality:
  ~(only US)~ GALLETTI, Bruno, c/o Novartis Vaccines and Diagnostics
    S.r.l., Via Fiorentina, 1, I-53100 Siena, IT Residence: IT Nationality:
  ~(only US)~ PARENTE, Pierino, c/o Novartis Vaccines and Diagnostics
    S.r.l., Via Fiorentina, 1, I-53100 Siena, IT Residence: IT Nationality:
  ~(only US)~ COSTANTINO, Paolo, c/o Novartis Vaccines and Diagnostics
    S.r.l., Via Fiorentina, 1, I-53100 Siena, IT Residence: IT Nationality:
Inventor: BERTI, Francesco, c/o Novartis Vaccines and Diagnostics S.r.l.,
    Via Fiorentina, 1, I-53100 Siena, IT Residence: IT Nationality: IT
  GALLETTI, Bruno, c/o Novartis Vaccines and Diagnostics S.r.l., Via
    Fiorentina, 1, I-53100 Siena, IT Residence: IT Nationality: IT
  PARENTE, Pierino, c/o Novartis Vaccines and Diagnostics S.r.l., Via
    Fiorentina, 1, I-53100 Siena, IT Residence: IT Nationality: IT
  COSTANTINO, Paolo, c/o Novartis Vaccines and Diagnostics S.r.l., Via
    Fiorentina, 1, I-53100 Siena, IT Residence: IT Nationality: IT
Agent: MARSHALL, Cameron, John, Carpmaels Ransford, 43-45 Bloomsbury
   Square, London WC1A 2RA, GB
Language: EN (33 pages, 3 drawings)
Application: WO 2007IB1855 A 20070321 (Local application)
Priority: GB 20065757 A 20060322
Designated States: (National Original) AE AG AL AM AT AU AZ BA BB BG BH BR
    BW BY BZ CA CH CN CO CR CU CZ DE DK DM DZ EC EE EG ES FI GB GD GE GH GM
    GT HN HR HU ID IL IN IS JP KE KG KM KN KP KR KZ LA LC LK LR LS LT LU LY
    MA MD MG MK MN MW MX MY MZ NA NG NI NO NZ OM PG PH PL PT RO RS RU SC SD
    SE SG SK SL SM SV SY TJ TM TN TR TT TZ UA UG US UZ VC VN ZA ZM ZW
  (Regional Original) AT BE BG BW CH CY CZ DE DK EA EE ES FI FR GB GH GM GR
    HU IE IS IT KE LS LT LU LV MC MT MW MZ NA NL OA PL PT RO SD SE SI SK SL
    SZ TR TZ UG ZM ZW
Original IPC: G01N-33/15(N,99,20060101,A,L) G01N-33/15(N,M,98,20060101,C)
    G01N-33/50(I,99,20060101,A,F) G01N-33/50(I,99,20060101,A,F)
    G01N-33/50(I.M.98,20060101,C)
Current IPC: G01N-33/15(N,99,20060101,A,L) G01N-33/15(N,M,98,20060101,C)
    G01N-33/50(I,99,20060101,A,F) G01N-33/50(I,99,20060101,A,F)
    G01N-33/50(I,M,98,20060101,C)
Current ECLA ICO class: K61K-39:60P10
Original Abstract: The invention is based on the use of a basic reagent
    under basic conditions to separate conjugated saccharide from
    unconjugated components in a sample, ~e.g.~ a vaccine, by precipitation
    of the conjugated saccharide. The invention allows rapid and
    quantitative separation of conjugated and conjugated components, which
   may be exploited in analytical methods for quantifying unconjugated
   saccharide or carrier. Therefore, the separation of conjugated and
    unconjugated components using the invention may be advantageously
    combined with a quantitative saccharide or carrier analysis to provide
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improved quality control for conjugate vaccines.

L'invention porte sur l'utilisation de reactifs basiques dans des conditions basiques pour separer des saccharides conjugues de saccharides non conjugues dans un echantillon, par exemple dans un vaccin, par precipitation du saccharide conjugue. L'invention permet une separation quantitative et rapide des composants conjugues d'avec les non conjugues, laquelle peut servir dans des methodes analytiques pour quantifier le saccharide non conjugues ou le vecteur. Ainsi, la separation de composants conjugues ou non conjugues peut se combiner avantageusement avec une analyse quantitative du saccharide ou du vecteur pour obtenir un controle de qualite ameliore des vaccins conjugues.

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Publication No. WO 2007122512 A3 (Update 200841 E)
Publication Date: 20080124
Language: EN
Designated States: (National Original) AE AG AL AM AT AU AZ BA BB BG BH BR
    BW BY BZ CA CH CN CO CR CU CZ DE DK DM DZ EC EE EG ES FI GB GD GE GH GM
    GT HN HR HU ID IL IN IS JP KE KG KM KN KP KR KZ LA LC LK LR LS LT LU LY
    MA MD MG MK MN MW MX MY MZ NA NG NI NO NZ OM PG PH PL PT RO RS RU SC SD
    SE SG SK SL SM SV SY TJ TM TN TR TT TZ UA UG US UZ VC VN ZA ZM ZW
  (Regional Original) AT BE BG BW CH CY CZ DE DK EA EE ES FI FR GB GH GM GR
    HU IE IS IT KE LS LT LU LV MC MT MW MZ NA NL OA PL PT RO SD SE SI SK SL
    SZ TR TZ UG ZM ZW
Original IPC: A61K-39/00(B, I, H, EP, 20060101, A, L)
    A61K-39/00(B,I,M,98,20060101,C) G01N-33/50(B,I,H,EP,20060101,A,F)
    G01N-33/50(B, I, M, 98, 20060101, C)
Current IPC: A61K-39/00(B,I,H,EP,20060101,20080124,A,L)
    A61K-39/00(B, I, H, EP, 20060101, 20080124, C, L)
    G01N-33/50(B, I, H, EP, 20060101, 20080124, A, F)
    G01N-33/50(B, I, H, EP, 20060101, 20080124, C, F)
Current ECLA ICO class: K61K-39:60P10
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10/7/11 (Item 11 from file: 351) DIALOG(R)File 351:Derwent WPI (c) 2011 Thomson Reuters. All rts. reserv.

0016568709 WPI ACC NO: 2007-283647/200727 XRAM Acc No: C2007-104024

Modifying bacterial ****capsular**** saccharide antigen comprises converting a neutral group in the saccharide into a cationic group, and/or converting a neutral group in the saccharide into an anionic group Patent Assignee: NOVARTIS VACCINES & DIAGNOSTICS INC (NOVS); NOVARTIS

VACCINES&DIAGNOSTICS INC (NOVS)
Inventor: BERTI F; TELFORD J; WACK A

Patent Family (5 patents, 116 countries)

Patent. Application Number Kind Date Kind Date Number Updat.e A 20060824 200727 WO 2007023386 A2 20070301 WO 2006IB2833 A 20060824 200845 E A 20060824 200857 E A 20060824 200857 E A 20060824 200917 E EP 1937304 A2 20080702 EP 2006831540 WO 2006IB2833 AU 2006283302 A1 20070301 AU 2006283302 CA 2620416 A1 20070301 CA 2620416 A 20060824 WO 2006IB2833 CA 2620416 A 20080225 US 20090136547 A1 20090528 WO 2006IB2833 A 20060824 200935 E US 200864663 A 20080825

Priority Applications (no., kind, date): GB 200517353 A 20050824; GB 20067738 A 20060419

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Patent Details
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Kind Lan Pg Dwg Filing Notes Number WO 2007023386 A2 EN 56 35

National Designated States, Original: AE AG AL AM AT AU AZ BA BB BG BR BW BY BZ CA CH CN CO CR CU CZ DE DK DM DZ EC EE EG ES FI GB GD GE GH GM HN HR HU ID IL IN IS JP KE KG KM KN KP KR KZ LA LC LK LR LS LT LU LV LY MA MD MG MK MN MW MX MY MZ NA NG NI NO NZ OM PG PH PL PT RO RS RU SC SD SE

SG SK SL SM SV SY TJ TM TN TR TT TZ UA UG US UZ VC VN ZA ZM ZW Regional Designated States, Original: AT BE BG BW CH CY CZ DE DK EA EE ES FI FR GB GH GM GR HU IE IS IT KE LS LT LU LV MC MW MZ NA NL OA PL PT RO

SD SE SI SK SL SZ TR TZ UG ZM ZW EP 1937304 A2 EN PCT Application WO 2006IB2833

Based on OPI patent WO 2007023386 Regional Designated States, Original: AT BE BG CH CY CZ DE DK EE ES FI FR

GB GR HU IE IS IT LI LT LU LV MC NL PL PT RO SE SI SK TR AU 2006283302 A1 EN Based on OPI patent WO 2007023386 PCT Application WO 2006IB2833 CA 2620416 A1 EN

PCT national entry CA 2620416 Based on OPI patent WO 2007023386 PCT Application WO 2006IB2833

US 20090136547 A1 EN

Alerting Abstract WO A2

NOVELTY - Modifying a bacterial ****capsular**** saccharide antigen comprises: (a) if the saccharide is anionic, converting a neutral group in the saccharide into a cationic group; (b) if the saccharide is cationic, converting a neutral group in the saccharide into an anionic group; (c) if the saccharide is neutral, converting a first neutral group in the saccharide into an anionic group and converting a second neutral group in the saccharide into a cationic group, thus providing a modified saccharide. DESCRIPTION - INDEPENDENT CLAIMS are:

- 1.a modified bacterial ****capsular**** saccharide, where the saccharide in its natural form includes repeating units that are cationic, but the saccharide in its modified form includes repeating units that are zwitterionic or anionic;
- 2.a modified bacterial ****capsular**** saccharide, where the saccharide in its natural form includes repeating units that are anionic, but the saccharide in its modified form includes repeating units that are zwitterionic or cationic;
- 3.a modified bacterial ****capsular**** saccharide, where the saccharide in its natural form includes repeating units that include either cationic or anionic groups, but the saccharide in its modified form includes repeating units that include both cationic and anionic groups; and
- 4.a modified bacterial ****capsular**** saccharide, where the saccharide includes a repeating unit that (a) includes both positively-charged and negatively-charged groups but (b) has no overall charge.

ACTIVITY - Antibacterial. No biological data given. MECHANISM OF ACTION - Vaccine.

USE - The method is useful for modifying a bacterial ****capsular**** saccharide antigen. The modified saccharide antigen can be used as active ingredients in compositions, e.g. vaccine, for the prevention and/or treatment of a bacterial infection, including diseases caused by ~S. agalactiae ~ , e.g. neonatal sepsis or bacteremia, neonatal pneumonia, neonatal meningitis, endometritis, osteomyelitis, or septic arthritis.

BIOTECHNOLOGY - Preferred Modified Saccharide: The repeating units in the modified saccharide are zwitterionic. It has both a free carboxyl group and a free amino group. Preferably, the saccharide is from group B streptococcus or meningococcus. However, the bacterial ****capsular**** saccharide is not from ~Bacillus fragilis ~ or ~Streptococcus pneumoniae ~ . A neutral group is converted to a group with a lower pKb value. An N-acetyl group is converted to an amino or amine group. Positive and negative charges are present on different monosaccharide within a repeating unit, where the positive and negative charges are not on adjacent monosaccharides within the repeating unit. At least 50% of the saccharide's repeating units are zwitterionic repeating units. The saccharide is a substantially full-length ****capsular******polysaccharide****. Preferred Method: In modifying a bacterial ****capsular**** saccharide antigen, deacetylating an N-acetyl group on the bacterial ****capsular**** saccharide in the presence of a base or enzyme to provide a free amino group. It further comprises reacting the free amino group with an aldehyde to provide an amine group, where the aldehyde is formaldehyde and the amine is a secondary amine. The N-acetyl group is present on a NeuAc moiety and/or a GlcNac moiety. The method also comprises reacting a carboxyl group on the bacterial ****capsular**** saccharide with pyruvate. It further comprises reacting the pyruvate with a carbodiimide or acetic acid. The method also comprises reacting a carboxyl group on the bacterial ****capsular**** saccharide with TEMPO (2,2,6,6-tetramethyl-1-piperidine oxoammonium ion) in the presence of hypochlorite and bromide. It also comprises hydrolysis of a terminal galactose unit on the bacterial ****capsular**** saccharide with O 3 /NO or beta-endogalactosidase. It also comprises oxidizing the terminal galactose unit with galactose oxidase to provide an aldehyde group the method also comprises reacting the aldehyde group with a free amino group or an amine group. It further comprises oxidizing NeuAc groups on the bacterial ****capsular**** saccharide to provide aldehyde groups and then reacting the aldehyde groups with a free amino group or an amine group.

Title Terms/Index Terms/Additional Words: MODIFIED; BACTERIA; CAPSULE; SACCHARIDE; ANTIGEN; COMPRISE; CONVERT; NEUTRAL; GROUP; CATION; ANION

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Class Codes
International Classification (+ Attributes)
IPC + Level Value Position Status Version
 A61K-0039/02 A I L B 20060101
 A61K-0039/09 A I F B 20060101
 A61K-0039/095 A I L B 20060101
 A61P-0031/04 A I L B 20060101
 C08B-0037/00 A I L B 20060101
 A61K S
                 20060101
 A61K-0039/02 C I L B 20090101
 A61K-0039/09 C I F B 20060101
 A61K-0039/09 C I B 20060101
 A61K-0039/095 C I L B 20090101
 A61P-0031/00 C I L B 20090101
 C08B-0037/00 C I L B 20060101
C08B-0037/00 C I B 20060101
ECLA: A61K-039/09A, A61K-039/095, C08B-037/00P
US Classification, Current Main: 424-244100; Secondary: 424-234100,
424-250100, 536-123100
US Classification, Issued: 424244.1, 536123.1, 424250.1, 424234.1
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File Segment: CPI
DWFI Class: B04; D16
Manual Codes (CPI/A-M): B04-B04C1; B04-C02F; B14-A01; B14-C09; B14-K01;
B14-N14; B14-N16; B14-S06; B14-S11B1; D05-A02; D05-H07

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Australia
Publication No. AU 2006283302 A1 (Update 200857 E)
Publication Date: 20070301
Assignee: NOVARTIS VACCINES & DIAGNOSTICS INC (NOVS)
Inventor: WACK A
  BERTI F
  TELFORD J
Language: EN
Application: AU 2006283302 A 20060824 (Local application)
Priority: GB 200517353 A 20050824
 GB 20067738 A 20060419
Related Publication: WO 2007023386 A (Based on OPI patent )
Original IPC: A61K-39/09(B,I,H,EP,20060101,20070705,A,F)
    C08B-37/00(B, I, H, EP, 20060101, 20070705, A, L)
Current IPC: A61K-39/09(B,I,H,EP,20060101,20070705,A,F)
    A61K-39/09(B, I, H, EP, 20060101, 20070705, C, F)
    C08B-37/00(B,I,H,EP,20060101,20070705,A,L)
    C08B-37/00(B,I,H,EP,20060101,20070705,C,L)
Current ECLA class: A61K-39/09A A61K-39/095
Canada
Publication No. CA 2620416 A1 (Update 200917 E)
Publication Date: 20070301
Assignee: NOVARTIS VACCINES&DIAGNOSTICS INC; IT (NOVS)
Inventor: BERTI F, IT
  TELFORD J. IT
 WACK A, IT
Language: EN
Application: CA 2620416 A 20060824 (Local application)
  WO 2006IB2833 A 20060824 (PCT Application)
  CA 2620416 A 20080225 (PCT national entry)
Priority: GB 200517353 A 20050824
  GB 20067738 A 20060419
Related Publication: WO 2007023386 A (Based on OPI patent )
Original IPC: A61K-39/09(B,I,H,EP,20060101,20070705,A,F)
    A61K-39/09(B, I, M, 98, 20060101, 20070705, C)
    C08B-37/00(B, I, H, EP, 20060101, 20070705, A, L)
    C08B-37/00(B, I, M, 98, 20060101, 20070705, C)
Current IPC: A61K-39/09(B,I,H,EP,20060101,20070705,A,F)
    A61K-39/09(B, I, M, 98, 20060101, 20070705, C)
    C08B-37/00(B, I, H, EP, 20060101, 20070705, A, L)
    C08B-37/00(B,I,M,98,20060101,20070705,C)
Current ECLA class: A61K-39/09A A61K-39/095 C08B-37/00P
Publication No. EP 1937304 A2 (Update 200845 E)
Publication Date: 20080702
**ZWITTERIONISATION KAPSELFORMIGER SACCHARIDE
  ZWITTERIONIZATION OF CAPSULAR SACCHARIDES
  ZWITTERIONISATION DE SACCHARIDES CAPSULAIRES**
Assignee: Novartis Vaccines and Diagnostics S.r.l., Via Fiorentina 1, 53100
    Siena (SI), IT
Inventor: TELFORD, John, Novartis Vaccines and Diagnostics S.r.l., Via
    Fiorentina, 1, I-53100 Siena, IT
  BERTI, Francesco, Novartis Vaccines and Diagnostics S.r.l., Via
    Fiorentina, 1, I-53100 Siena, IT
  WACK, Andreas, Novartis Vaccines and Diagnostics S.r.l., Via Fiorentina,
    1, I-53100 Siena, IT
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Agent: Marshall, Cameron John, Carpmaels Ransford, 43-45 Bloomsbury
    Square, London WC1A 2RA, GB
Language: EN
Application: EP 2006831540 A 20060824 (Local application)
  WO 2006IB2833 A 20060824 (PCT Application)
Priority: GB 200517353 A 20050824
  GB 20067738 A 20060419
Related Publication: WO 2007023386 A (Based on OPI patent )
Designated States: (Regional Original) AT BE BG CH CY CZ DE DK EE ES FI FR
    GB GR HU IE IS IT LI LT LU LV MC NL PL PT RO SE SI SK TR
Original IPC: A61K-39/09(B,I,H,EP,20060101,20080327,A,F)
    A61K-39/09(B, I, M, 98, 20060101, 20080327, C)
    C08B-37/00(B, I, H, EP, 20060101, 20080327, A, L)
    C08B-37/00(B,I,M,98,20060101,20080327,C)
Current IPC: A61K-39/09(B,I,H,EP,20060101,20080327,A,F)
    A61K-39/09(B, I, H, EP, 20060101, 20080327, C, F)
    C08B-37/00(B, I, H, EP, 20060101, 20080327, A, L)
    C08B-37/00(B,I,H,EP,20060101,20080327,C,L)
Current ECLA class: A61K-39/09A A61K-39/095 C08B-37/00P
Original Abstract: Capsular saccharides are typically anionic. In the
    invention, however, cationic groups are introduced, such that the
    modified saccharide has a repeating unit which includes both cationic
    and anionic groups. These cationic and anionic groups can be balanced
    to give a zwitterionic repeating unit. These modifications can convert
    a saccharide that is normally a T-independent antigen into one that can
    activate T cells without requiring conjugation to a carrier. Typically,
    the invention modifies an anionic bacterial capsular saccharide antigen
    by converting a neutral group in the saccharide into a cationic group
    e.g. to change -NHAc to -NH3+.
United States
Publication No. US 20090136547 A1 (Update 200935 E)
Publication Date: 20090528
**ZWITTERIONIZATION OF CAPSULAR SACCHARIDES**
Assignee: NOVARTIS VACCINES AND DIAGNOSTICS SRL, Siena, IT (NOVS)
 Telford, John, Siena, IT Residence: IT
  Berti, Francesco, Siena, IT Residence: IT
  Wack, Andreas, Siena, IT Residence: IT
Inventor: Berti, Francesco, Siena, IT Residence: IT
  Telford, John, Siena, IT Residence: IT
  Wack, Andreas, Siena, IT Residence: IT
Agent: NOVARTIS VACCINES AND DIAGNOSTICS INC., INTELLECTUAL PROPERTY R338,
    P.O. BOX 8097, Emeryville, CA, US
Language: EN
Application: US 200864663 A 20080825 (Local application)
  WO 2006IB2833 A 20060824 (PCT Application)
Priority: GB 200517353 A 20050824
  GB 20067738 A 20060419
Original IPC: A61K-39/02(B,I,H,US,20060101,20090528,A,L)
    A61K-39/02(B, I, M, 98, 20060101, 20090528, C)
    A61K-39/09(B, I, H, US, 20060101, 20090528, A, F)
    A61K-39/09(B, I, M, 98, 20060101, 20090528, C)
    A61K-39/095(B, I, H, US, 20060101, 20090528, A, L)
    A61K-39/095(B,I,M,98,20060101,20090528,C)
    A61P-31/00(B, I, M, 98, 20060101, 20090528, C)
   A61P-31/04(B, I, H, US, 20060101, 20090528, A, L)
   C08B-37/00(B,I,H,US,20060101,20090528,A,L)
    C08B-37/00(B, I, M, 98, 20060101, 20090528, C)
Current IPC: A61K-39/02(B,I,H,US,20060101,20090528,A,L)
    A61K-39/02(B,I,H,US,20090101,20090528,C,L)
    A61K-39/09(B, I, H, US, 20060101, 20090528, A, F)
    A61K-39/09(B, I, H, US, 20090101, 20090528, C, F)
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A61K-39/095(B, I, H, US, 20060101, 20090528, A, L)
    A61K-39/095(B, I, H, US, 20090101, 20090528, C, L)
   A61P-31/00(B, I, H, US, 20090101, 20090528, C, L)
    A61P-31/04(B, I, H, US, 20060101, 20090528, A, L)
    C08B-37/00(B, I, H, US, 20060101, 20090528, A, L)
    C08B-37/00(B, I, H, US, 20090101, 20090528, C, L)
Current ECLA class: A61K-39/09A A61K-39/095 C08B-37/00P
Current US Class (main): 424-244100
Current US Class (secondary): 424-234100 424-250100 536-123100
Original US Class (main): 424244.1
Original US Class (secondary): 536123.1 424250.1 424234.1
Original Abstract: Capsular saccharides are typically anionic. In the
    invention, however, cationic groups are introduced, such that the
    modified saccharide has a repeating unit which includes both cationic
    and anionic groups. These cationic and anionic groups can be balanced
    to give a zwitterionic repeating unit. These modifications can convert
    a saccharide that is normally a T-independent antigen into one that can
    activate T cells without requiring conjugation to a carrier. Typically,
    the invention modifies an anionic bacterial capsular saccharide antigen
    by converting a neutral group in the saccharide into a cationic group
    e.g. to change --NHAc to --NH3 +.
Claim:
**1**. A method for modifying a bacterial capsular saccharide
        antigen, comprising a step of:
    * (i) if the saccharide is anionic, converting a neutral group in the
          saccharide into a cationic group;
    * (ii) if the saccharide is cationic, converting a neutral group in the
          saccharide into an anionic group;
    * (iii) if the saccharide is neutral, converting a first neutral group
          in the saccharide into an anionic group and converting a second
          neutral group in the saccharide into a cationic group,
  thereby
          providing a modified saccharide.
Publication No. WO 2007023386 A2 (Update 200727 B)
Publication Date: 20070301
**ZWITTERIONIZATION OF CAPSULAR SACCHARIDES
  ZWITTERIONISATION DE SACCHARIDES CAPSULAIRES**
Assignee: ~ (except US) ~ NOVARTIS VACCINES AND DIAGNOSTICS SRL, Via
    Fiorentina 1, I-53100 Siena, IT Residence: IT Nationality: IT (NOVS)
  ~(only US)~ TELFORD, John, Novartis Vaccines and Diagnostics S.r.l., Via
    Fiorentina, 1, I-53100 Siena, IT Residence: IT Nationality: IT
  ~(only US)~ BERTI, Francesco, Novartis Vaccines and Diagnostics S.r.l.,
    Via Fiorentina, 1, I-53100 Siena, IT Residence: IT Nationality: IT
  ~(only US)~ WACK, Andreas, Novartis Vaccines and Diagnostics S.r.l., Via
    Fiorentina, 1, I-53100 Siena, IT Residence: IT Nationality: DE
Inventor: TELFORD, John, Novartis Vaccines and Diagnostics S.r.1., Via
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Language: EN (56 pages, 35 drawings)
Application: WO 2006IB2833 A 20060824 (Local application)
Priority: GB 200517353 A 20050824
 GB 20067738 A 20060419
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Designated States: (National Original) AE AG AL AM AT AU AZ BA BB BG BR BW BY BZ CA CH CN CO CR CU CZ DE DK DM DZ EC EE EG ES FI GB GD GE GH GM HN HR HU ID IL IN IS JP KE KG KM KN KP KR KZ LA LC LK LR LS LT LU LV LY AM MD MG MK NN MW MX MY MZ NA NG NI NO NZ OM PG PH PL PT RO RS RU SC SD SE SG SK SL SM SV SY TJ TM TN TR TT TZ UA UG US UZ VC VN ZA ZM ZW (Regional Original) AT BE BG BW CH CY CZ DE DK EA EE ES FI FR GB GH GM GR

(Regional Original) Al BE BG BW CH CI CZ DE DK EA EE ES FI FR GB GH GW GK U IE IS IT KE LS LT LU LV MC MW MZ NA NL OA PL PT RO SD SE SI SK SL SZ TR TZ UG ZM ZW

Original IPC: A61K(99,20060101,S)

Current IPC: A61K(99,20060101,S)

Current ECLA class: A61K-39/09A A61K-39/095 C08B-37/00P

Original Abstract: Capsular saccharides are typically anionic. In the invention, however, cationic groups are introduced, such that the modified saccharide has a repeating unit which includes both cationic and anionic groups. These cationic and anionic groups can be balanced to give a zwitterionic repeating unit. These modifications can convert a saccharide that is normally a T-independent antigen into one that can activate T cells without requiring conjugation to a carrier. Typically, the invention modifies an anionic bacterial capsular saccharide antigen by converting a neutral group in the saccharide into a cationic group e.g. to change -NHAC to -NH3 +.

Les saccharides capsulaires sont typiquement anioniques. Neanmoins dans l'invention on introduit des groupes cationiques pour que le saccharide modifie comporte une unite repetitive incluant a la fois des groupes cationiques et anioniques. Ces groupes anioniques et cationiques peuvent etre equilibres pour donner une unite zwitterionique repetitive. Ces modifications peuvent convertir un saccharide qui est normalement un antigene al independant en un antigene activateur de cellules T sans necessiter de le conjuguer a un porteur. Typiquement, l'invention modifie un antigene anionique bacterien de saccharide capsulaire en convertissant un groupe neutre du saccharide en un groupe cationique, par exemple en changeant -NHAC en -NHA3 +.

10/7/12 (Item 12 from file: 351) DIALOG(R)File 351:Derwent WPI (c) 2011 Thomson Reuters. All rts. reserv.

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WPI ACC NO: 2006-559893/200657

Preparing conjugate of Streptococcus agalactiae ****capsular**** saccharide and carrier molecule comprises e.g. either oxidizing the saccharide, reductive amination, producing activated saccharide and reacting with carrier molecule

Patent Assignee: CHIRON SRL (CHIR); NOVARTIS VACCINES & DIAGNOSTICS INC (NOVS); BERTI F (BERT-1); NOVARTIS VACCINES&DIAGNOSTICS INC (NOVS) Inventor: BERTI F; FRANCESCO B

Patent Family (10 patents, 112 countries)

Patent						Application				
	Number		Kind	Date	Nur	nber	Kind	Date	Update	
	WO	2006082530	A2	20060810	WO	2006IB756	A	20060201	200657	В
	EP	1846038	A2	20071024	EP	2006727404	A	20060201	200771	E
					WO	2006IB756	A	20060201		
	AU	2006211052	A1	20060810	AU	2006211052	A	20060201	200780	E
	WO	2006082530	A3	20080703					200845	E
	JP	2008532930	W	20080821	WO	2006IB756	A	20060201	200857	E
					JP	2007553744	A	20060201		
	MX	2007009277	A1	20070901	WO	2006IB756	A	20060201	200864	E
					MX	20079277	A	20070801		
	US	20090043077	A1	20090212	WO	2006IB756	A	20060201	200912	E
					US	2008883614	A	20080318		
	CN	101304765	A	20081112	CN	200680007342	2 A	20060201	200918	E

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WO 2006IB756 A 20060201
                 20090128 ZA 20076969
                                       A 20070820 200918 E
ZA 200706969
             A
                                        A 20060201 201106 E
NZ 560432
                 20101224 NZ 560432
             A
                          WO 2006IB756
                                        A 20060201
Priority Applications (no., kind, date): GB 20052095 A 20050201
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Patent Details

Number Kind Lan Pg Dwg Filing Notes 48 WO 2006082530 A2 EN

National Designated States, Original: AE AG AL AM AT AU AZ BA BB BG BR BW BY BZ CA CH CN CO CR CU CZ DE DK DM DZ EC EE EG ES FI GB GD GE GH GM HR HU ID IL IN IS JP KE KG KM KN KP KR KZ LC LK LR LS LT LU LV LY MA MD MG MK MN MW MX MZ NA NG NI NO NZ OM PG PH PL PT RO RU SC SD SE SG SK SL SM SY TJ TM TN TR TT TZ UA UG US UZ VC VN YU ZA ZM ZW

Regional Designated States, Original: AT BE BG BW CH CY CZ DE DK EA EE ES

FI FR GB GH GM GR HU IE IS IT KE LS LT LU LV MC MW MZ NA NL OA PL PT RO SD SE SI SK SL SZ TR TZ UG ZM ZW A2 EN EP 1846038 PCT Application WO 2006IB756

Based on OPI patent WO 2006082530 Regional Designated States, Original: AL AT BA BE BG CH CY CZ DE DK EE ES FI FR GB GR HR HU IE IS IT LI LT LU LV MC MK NL PL PT RO SE SI SK TR YU AU 2006211052 A1 EN Based on OPI patent WO 2006082530 A3 EN WO 2006082530

National Designated States, Original: AE AG AL AM AT AU AZ BA BB BG BR BW BY BZ CA CH CN CO CR CU CZ DE DK DM DZ EC EE EG ES FI GB GD GE GH GM HR HU ID IL IN IS JP KE KG KM KN KP KR KZ LC LK LR LS LT LU LV LY MA MD MG MK MN MW MX MZ NA NG NI NO NZ OM PG PH PL PT RO RU SC SD SE SG SK SL SM SY TJ TM TN TR TT TZ UA UG US UZ VC VN YU ZA ZM ZW

Regional Designated States, Original: AT BE BG BW CH CY CZ DE DK EA EE ES FI FR GB GH GM GR HU IE IS IT KE LS LT LU LV MC MW MZ NA NL OA PL PT RO

SD SE SI SK SL SZ TR TZ UG ZM ZW JP 2008532930 W JA 43

> А EN

MX 2007009277 A1 ES US 20090043077 A1 EN CN 101304765 zHZA 200706969 A EN PCT Application WO 2006IB756 Based on OPI patent WO 2006082530 PCT Application WO 2006IB756 Based on OPI patent WO 2006082530 PCT Application WO 2006IB756 PCT Application WO 2006IB756 Based on OPI patent WO 2006082530

PCT Application WO 2006IB756 Based on OPI patent WO 2006082530

Alerting Abstract WO A2

NZ 560432

NOVELTY - Preparing a conjugate of a ~Streptococcus agalactiae ~****capsular**** saccharide (I) and a carrier molecule, comprises: either oxidizing (I) to introduce aldehyde group, reductive amination of the aldehyde group, producing activated saccharide or de-N-acetylating (I), reacting the saccharide with bifunctional linker and reacting activated saccharide with a carrier molecule; or oxidizing (I) to introduce an aldehyde group into a galactose residue in the saccharide; and coupling to a carrier.

DESCRIPTION - Preparing a conjugate of a ~Streptococcus agalactiae ~ ****capsular**** saccharide (I) and a carrier molecule, comprises: either oxidizing (I) to introduce an aldehyde group into at least one terminal sialic acid residue in the saccharide, reductive amination of the aldehyde group with ammonia or a primary amine, to give a methylamine; reacting the methylamine with a bifunctional linker, to give an activated saccharide; and reacting the activated saccharide with a carrier molecule; or de-N-acetylating (I) to give a de-N-acetylated saccharide; reacting the de-N-acetylated saccharide with a bifunctional linker to give an activated saccharide; and reacting the activated saccharide with a carrier molecule; or oxidizing (I) to introduce an aldehyde group into at least one galactose residue in the saccharide, to give a modified galactose residue; and coupling the modified galactose residue to a carrier molecule.

An INDEPENDENT CLAIM is also included for the conjugate, comprising (I) moiety joined to a carrier via a linker moiety, other the linker moiety is attached to a sialic acid residue in the ****capsular**** saccharide moiety.

ACTIVITY - Antibacterial.

No biological data given.

MECHANISM OF ACTION - Vaccine.

USE - The conjugate is useful for immunization.

ADVANTAGE - The process retains the sialic acid residues in a form of closure than the native ****polysaccharide**** and improves the coupling properties.

Technology Focus

BIOTECHNÓLOGY - Preferred Process: The saccharide is chemically modified relative to the native ****capsular**** saccharide and is de-O-acetylated or de-N-acetylated (partially or fully). The aldehyde groups are introduced into 5-50% of the total sialic acid monosaccharide units. After conjugation, free and conjugated saccharides are separated. The oxidizing step introduces the aldehyde chemically and involves the use of a periodate salt to oxidize vicinal hydroxides. The reductive amination involves either ammonia or a primary amine of formula (NH 2 R), preferably an ammonium salt in combination with a reducing agent. The reactions with both the saccharide and the carrier involve amines, and where the linker has formula X-L-X, where the two X groups are N-oxysuccinimide and can react with the amines; and L is a linking moiety in the linker (adipic acid N-hydroxysuccinimide diester). The saccharide is, if necessary, substantially totally re-N-acetylated prior to reductive amination. The

substantially totally re-M-acetylated prior to reductive amination. The individual saccharide is attached to multiple carriers.

Preferred Components: In the conjugate, the saccharide is from one of GBS

serotypes Ia, Ib, II, III or V and has its native form. The saccharide is shorter than the native *****capsular***** saccharide. The carrier is diphtheria toxoid, tetanus toxoid, CRM197, human serum albumin, an artificial protein comprising multiple human CD4* I cell epitopes from various pathogen-derived antigens, protein D from -H. influenzae ~, or a ~S. agalactiae ~ protein. The carrier is attached to the saccharide via an amine group in the carrier. The conjugate has a saccharide:protein ratio (w/w) of 1:5-5:1. The bifunctional linker is hetero or homo-bifunctional

Title Terms/Index Terms/Additional Words: PREPARATION; CONJUGATE; STREPTCOCCCUS; CAPSULE; SACCHARIDE; CARRY; MOLECULAR; COMPRISE; OXIDATION , REDUCE; AMINATE; PRODUCE; ACTIVATE; REACT Class Codes

International Classification (+ Attributes)

IPC + Level Value Position Status Version A61K-0039/09 A I R 20060101
A61K-039/09 A I F B 20060101
A61K-039/09 A I L B 20060101
A61K-039/09 A I L B 20060101
A61K-039/395 A I L B 20060101
A61K-0039/395 A I L B 20060101
A61K-0047/48 A I F 20060101
A61K-0047/48 A I F 20060101
A61K-0047/48 A I F B 20060101
A61K-0047/48 A I L B 20060101
A61K-0037/04 A I L B 20060101

C07K-0001/00 A I L B 20060101 C07K-0014/765 A I F B 20060101 C08B-0037/00 A I L B 20060101

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A61P-0039/04 A I L
                          20060101
 A61K S I B 20090101
 A61K-0039/09 C I
                          20060101
 A61K-0039/09 C I
                      R 20060101
 A61K-0039/09 C I F B 20060101
 A61K-0039/09 C I L B 20060101
 A61K-0039/385 C I L B 20060101
 A61K-0047/48 C I
                          20060101
 A61K-0047/48 C I R 20060101
 A61K-0047/48 C I F B 20060101
 A61P-0031/00 C I L B 20060101
 A61P-0037/00 C I
                          20060101
 A61P-0037/00 C I L B 20060101
 A61P-0039/00 C I L B 20060101
 C07K-0001/00 C I L B 20090101
 C07K-0014/435 C I F B 20090101
 C08B-0037/00 C I L B 20090101
ECLA: A61K-039/09, A61K-047/48R2D, A61K-047/48R2L, A61K-047/48R2V
ICO: K61K-039:60P10
US Classification, Current Main: 530-363000; Secondary: 530-402000,
536-124000
US Classification, Issued: 530363, 536124, 530402
JP Classification
 FI Term
                  Facet Rank Type
A61K-039/09
A61K-039/385
A61P-031/04
A61P-037/04
F-Term View Point Additional
Theme
      + Figure Code
4C085
4C201
4C085
          AA03
4C085
         BA14
4C085
         BA38
4C085
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4C085
         CC24
4C085
         DD59
4C085
         DD62
4C085
         EE06
40085
         FF24
File Segment: CPI
DWPI Class: A11; A96; B04; D16
Manual Codes (CPI/A-M): A03-A01; A12-V01; B04-B04C; B04-C02F; B04-N02;
 B04-N03; B14-A01; B14-S11B1; D05-H07
Original Publication Data by Authority
Australia
Publication No. AU 2006211052 A1 (Update 200780 E)
Publication Date: 20060810
Assignee: NOVARTIS VACCINES & DIAGNOSTICS INC (NOVS)
Inventor: BERTI F
Language: EN
Application: AU 2006211052 A 20060201 (Local application)
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Priority: GB 20052095 A 20050201
Related Publication: WO 2006082530 A (Based on OPI patent )
Original IPC: A61K-47/48(B,I,H,EP,20060101,20060915,A,F)
Current IPC: A61K-39/09(R,I,M,EP,20060101,20080531,A)
    A61K-39/09(R, I, M, EP, 20060101, 20080531, C)
    A61K-47/48(B, I, H, EP, 20060101, 20060915, A, F)
    A61K-47/48(B, I, H, EP, 20060101, 20060915, C, F)
Current ECLA class: A61K-39/09 A61K-47/48R2D A61K-47/48R2L A61K-47/48R2V
Current ECLA ICO class: K61K-39:60P10
China
Publication No. CN 101304765 A (Update 200918 E)
Publication Date: 20081112
**Conjugation of streptococcal capsular saccharides**
Assignee: NOVARTIS VACCINES DIAGNOSTICS INC; IT (NOVS)
Inventor: FRANCESCO, BERTI, IT
Language: ZH
Application: CN 200680007342 A 20060201 (Local application)
  WO 2006IB756 A 20060201 (PCT Application)
Priority: GB 20052095 A 20050201
Related Publication: WO 2006082530 A (Based on OPI patent )
Original IPC: A61K-39/09(I.CN, 20060101, A.L) A61K-39/09(I.M, 98, 20060101, C)
    A61K-47/48(I,CN,20060101,A,F) A61K-47/48(I,M,98,20060101,C)
    A61P-37/00(I,M,98,20060101,C) A61P-37/04(I,CN,20060101,A,L)
Current IPC: A61K-39/09(A,I,CN,20060101,A,L) A61K-39/09(I,M,98,20060101,C)
    A61K-47/48(I,CN,20060101,A,F) A61K-47/48(I,M,98,20060101,C)
    A61P-37/00(I,M,98,20060101,C) A61P-37/04(I,CN,20060101,A,L)
Current ECLA class: A61K-39/09 A61K-47/48R2D A61K-47/48R2L A61K-47/48R2V
Current ECLA ICO class: K61K-39:60P10
Original Abstract: This invention claims three conjugation methods for use
    with the capsular saccharide of Streptococcus agalactiae. In the first
    method, reductive animation of oxidised sialic acid residue side chains
    is used, but the aldehyde groups are first aminated, and then the amine
    is coupled to a carrier via a linker. In the second method, sialic acid
    residues and/or N-acetyl-glucosamine residues are de-N-acetylated to
    give amine groups, and the amine groups are coupled to a carrier
    protein via a linker. In the third method, linkage is via galactose
    residues in the capsular saccharide rather than sialic acid residues,
    which can conveniently be achieved using galactose oxidase.
Claim: [CLAIM 1] A process for preparing a conjugate of a Streptococcus
    agalactiae capsular saccharide and a carrier molecule, comprising the
    steps of: (a) oxidising a S.agalactiae capsular saccharide in order to
    introduce an aldehyde group into at least one terminal sialic acid
    residue in the saccharide; (b) subjecting the aldehyde group to
    reductive amination with ammonia or a primary amine, to give a
    -CH2-linked amine; (c) reacting the -CH2-linked amine with a
    bifunctional linker, to give an activated saccharide; and (d) reacting
    the activated saccharide with a carrier molecule, thereby giving the
    conjugate.
  [CLAIM 2] A conjugate, comprising a Streptococcus agalactiae capsular
    saccharide moiety joined to a carrier via a linker moiety, wherein the
    linker moiety is attached to a sialic acid residue in the capsular
    saccharide moietv.
  [CLAIM 3] The conjugate according to claim 2, obtainable by the process
    according to claim 1.
  [CLAIM 4] The conjugate or process according to any preceding claim,
    wherein the saccharide is from one of GBS serotypes Ia, Ib, II, III or
  [CLAIM 5] The conjugate or process according to any preceding claim,
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wherein the saccharide has its native form. [CLAIM 6] The conjugate or process according to any one of claims 1 to 4, wherein the saccharide is shorter than the native capsular saccharide.

- [CLAIM 7] The conjugate or process according to any one of claims 1 to 4, wherein the saccharide is chemically modified relative to the native capsular saccharide.
- [CLAIM 8] The conjugate or process according to claim 7, wherein the saccharide is de-O-acetylated partially or fully.
- [CLAIM 9] The conjugate or process according to claim 7, wherein the saccharide is de-N-acetylated partially or fully.
- [CLAIM 10] The conjugate or process according to any preceding claim, wherein the carrier is diphtheria toxoid, tetanus toxoid, CRM 197, human serum albumin, an artificial protein comprising multiple human CD4+T cell epitopes from various pathogen-derived antigens, protein D from H. influenzae, or a S.aqulactiae protein.
- [CLAIM 11] The conjugate or process according to any preceding claim, wherein the carrier is attached to the saccharide via a -NH2 group in the carrier.
- [CLAIM 12] The conjugate or process according to any preceding claim, wherein the conjugate has a saccharide: protein ratio (w/w) of between 1:5 and 5:1.
- [CLAIM 13] The conjugate or process according to any preceding claim, wherein aldehyde groups are introduced into between 5% and 50% of the total stalic acid monosaccharide units.
- [CLAIM 14] The process according to any preceding claim, wherein, after conjugation, free and conjugated saccharides are separated.
- [CLAIM 15] The process according to any preceding claim, wherein step (a) introduces the aldehyde chemically.
- [CLAIM 16] The process according to claim 15, wherein step (a) involves the use of a periodate salt to oxidise vicinal hydroxides.
- [CLAIM 17] The process according to any preceding claim, wherein
- reductive amination involves either ammonia or a primary amine (NH2R). [CLAIM 18] The process according to claim 17, wherein reductive amination
- involves an ammonium salt in combination with a reducing agent. [CLAIM 19] The process according to any preceding claim, wherein the
- bifunctional linker is heterobifunctional. [CLAIM 20] The process according to any one of claims 1 to 18, wherein
- the bifunctional linker is homobifunctional. [CLAIM 21] The process according to claim 20, where the reactions with
- both the saccharide and the carrier involve amines, and wherein the linker has formula X-L-X, wherein: the two X groups are the same as each other and can react with the amines; and L is a linking moiety in the linker.
- [CLAIM 22] The process according to claim 21, wherein X is N-oxysuccinimide.
- [CLAIM 23] The process according to claim 22, wherein the linker is adipic acid N-hydroxysuccinimide diester.
- [CLAIM 24] The process according to any preceding claim, wherein the saccharide is, if necessary, substantially totally re-N-acetylated prior to reductive amination.
- [CLAIM 25] The process or conjugate according to any preceding claim, wherein an individual saccharide is attached to multiple carriers.
- [CLAIM 26] A process for preparing a conjugate of a Streptococcus agalactiae capsular saccharide and a carrier molecule, comprising the steps of: (a) de-N-acetylating the capsular saccharide, to give a de-N-acetylated saccharide; (b) reacting the de-N-acetylated saccharide with a bifunctional linker, to give an activated saccharide; and (c) reacting the activated saccharide with a carrier molecule, thereby giving the conjugate.
- [CLAIM 27] A process for preparing a conjugate of a capsular saccharide and a carrier molecule, comprising the steps of: (a) oxidising a capsular saccharide in order to introduce an aldehyde group into at least one galactose residue in the saccharide, to give a modified galactose residue; and (b) coupling the modified galactose residue to a carrier molecule.

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EPO
Publication No. EP 1846038 A2 (Update 200771 E)
Publication Date: 20071024
**KONJUGATION VON STREPTOKOKKEN-KAPSELSACCHARIDEN
  CONJUGATION OF STREPTOCOCCAL CAPSULAR SACCHARIDES
  CONJUGAISON DE SACCHARIDES CAPSULAIRES STREPTOCOCCIQUES**
Assignee: Novartis Vaccines and Diagnostics S.r.l., Via Fiorentina 1, 53100
    Siena (SI), IT (NOVS)
Inventor: BERTI, Francesco, Chiron Vaccines, Via Fiorentina, 1, I-53100
    Siena, IT
Agent: Marshall, Cameron John, Carpmaels Ransford, 43-45 Bloomsbury
    Square, London WC1A 2RA, GB
Language: EN
Application: EP 2006727404 A 20060201 (Local application)
  WO 2006IB756 A 20060201 (PCT Application)
Priority: GB 20052095 A 20050201
Related Publication: WO 2006082530 A (Based on OPI patent )
Designated States: (Regional Original) AL AT BA BE BG CH CY CZ DE DK EE ES
    FI FR GB GR HR HU IE IS IT LI LT LU LV MC MK NL PL PT RO SE SI SK TR YU
Original IPC: A61K-47/48(B,I,H,EP,20060101,20060814,A,F)
    A61K-47/48(B, I, M, 98, 20060101, 20060814, C)
Current IPC: A61K-39/09(B,I,H,EP,20060101,20080905,A,L)
    A61K-39/09(B, I, H, EP, 20060101, 20080905, C, L)
    A61K-47/48(B, I, H, EP, 20060101, 20080905, A, F)
    A61K-47/48(B, I, H, EP, 20060101, 20080905, C, F)
    A61P-39/00(B, I, H, EP, 20060101, 20080905, C, L)
    A61P-39/04(B, I, H, EP, 20060101, 20080905, A, L)
Current ECLA class: A61K-39/09 A61K-47/48R2D A61K-47/48R2L A61K-47/48R2V
Current ECLA ICO class: K61K-39:60P10
Original Abstract: Three conjugation methods for use with the capsular
    saccharide of Streptococcus agalactiae. In the first method, reductive
    animation of oxidised sialic acid residue side chains is used, but the
    aldehyde groups are first aminated, and then the amine is coupled to a
    carrier via a linker. In the second method, sialic acid residues and/or
    N-acetyl-glucosamine residues are de-N-acetylated to give amine groups,
    and the amine groups are coupled to a carrier protein via a linker. In
    the third method, linkage is via galactose residues in the capsular
    saccharide rather than sialic acid residues, which can conveniently be
    achieved using galactose oxidase.
Japan
Publication No. JP 2008532930 W (Update 200857 E)
Publication Date: 20080821
Language: JA (43 pages)
Application: JP 2007553744 A 20060201 (Local application)
  WO 2006IB756 A 20060201 (PCT Application)
Priority: GB 20052095 A 20050201
Related Publication: WO 2006082530 A (Based on OPI patent )
Original IPC: A61K-39/09(B,I,H,JP,20060101,20080725,A,F)
    A61K-39/09(B, I, M, 98, 20060101, 20080725, C)
    A61K-39/385 (B, I, H, JP, 20060101, 20080725, A, L)
    A61K-39/385(B, I, M, 98, 20060101, 20080725, C)
    A61P-31/00(B, I, M, 98, 20060101, 20080725, C)
    A61P-31/04(B, I, H, JP, 20060101, 20080725, A, L)
    A61P-37/00(B, I, M, 98, 20060101, 20080725, C)
    A61P-37/04(B,I,H,JP,20060101,20080725,A,L)
Current IPC: A61K-39/09(B,I,H,JP,20060101,20080725,A,F)
   A61K-39/09(B,I,H,JP,20060101,20080725,C,F)
    A61K-39/385(B, I, H, JP, 20060101, 20080725, A, L)
    A61K-39/385(B, I, H, JP, 20060101, 20080725, C, L)
    A61K-47/48(R, I, M, EP, 20060101, 20060722, A)
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A61K-47/48(R, I, M, EP, 20060101, 20060722, C)
    A61P-31/00(B, I, H, JP, 20060101, 20080725, C, L)
   A61P-31/04(B,I,H,JP,20060101,20080725,A,L)
    A61P-37/00(B, I, H, JP, 20060101, 20080725, C, L)
    A61P-37/04(B, I, H, JP, 20060101, 20080725, A, L)
Current ECLA class: A61K-39/09 A61K-47/48R2D A61K-47/48R2L A61K-47/48R2V
Current ECLA ICO class: K61K-39:60P10
Current JP F-Terms: 4C085 4C201 4C085AA03 4C085BA14 4C085BA38 4C085BB11
    4C085BB15 4C085BB24 4C085CC07 4C085CC24 4C085DD59 4C085DD62 4C085EE06
    4C085FF24
Mevico
Publication No. MX 2007009277 A1 (Update 200864 E)
Publication Date: 20070901
Assignee: NOVARTIS VACCINES & DIAGNOSTICS INC (NOVS)
Inventor: BERTI F
Language: ES
Application: WO 2006IB756 A 20060201 (PCT Application)
 MX 20079277 A 20070801 (Local application)
Priority: GB 20052095 A 20050201
Related Publication: WO 2006082530 A (Based on OPI patent )
Original IPC: A61K-47/48(I,MX,20060101,A,F) A61K-47/48(I,M,98,20060101,C)
Current IPC: A61K-47/48(I,MX,20060101,A,F) A61K-47/48(I,M,98,20060101,C)
New Zealand
Publication No. NZ 560432 A (Update 201106 E)
Publication Date: 20101224
Assignee: NOVARTIS VACCINES&DIAGNOSTICS INC (NOVS)
Inventor: BERTI F
Language: EN
Application: NZ 560432 A 20060201 (Local application)
  WO 2006IB756 A 20060201 (PCT Application)
Priority: GB 20052095 A 20050201
Related Publication: WO 2006082530 A (Based on OPI patent )
Original IPC: A61K-39/09(I,NZ,20060101,A,L) A61K-47/48(I,NZ,20060101,A,F)
   A61P-39/04(I,NZ,20060101,A,L)
Current IPC: A61K-39/09(I,NZ,20060101,A,L) A61K-47/48(I,NZ,20060101,A,F)
    A61P-39/04(I,NZ,20060101,A,L)
United States
Publication No. US 20090043077 A1 (Update 200912 E)
Publication Date: 20090212
**Conjugation of streptococcal capsular saccharides**
Assignee: Berti, Francesco, Siena, IT Residence: IT (BERT-I)
Inventor: Berti, Francesco, Siena, IT Residence: IT
Agent: NOVARTIS VACCINES AND DIAGNOSTICS INC., INTELLECTUAL PROPERTY R338,
    P.O. BOX 8097, Emeryville, CA, US
Language: EN
Application: US 2008883614 A 20080318 (Local application)
  WO 2006IB756 A 20060201 (PCT Application)
Priority: GB 20052095 A 20050201
Original IPC: C07K-1/00(B,I,H,US,20060101,20090212,A,L)
    C07K-1/00(B,I,M,98,20060101,20090212,C)
    C07K-14/435(B, I, M, 98, 20060101, 20090212, C)
    C07K-14/765(B, I, H, US, 20060101, 20090212, A, F)
    C08B-37/00(B, I, H, US, 20060101, 20090212, A, L)
    C08B-37/00(B,I,M,98,20060101,20090212,C)
Current IPC: A61K-39/09(R,I,M,EP,20060101,20080531,A)
    A61K-39/09(R, I, M, EP, 20060101, 20080531, C)
    A61K-47/48(R, I, M, EP, 20060101, 20060722, A)
    A61K-47/48(R, I, M, EP, 20060101, 20060722, C)
    C07K-1/00(B, I, H, US, 20060101, 20090212, A, L)
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C07K-1/00(B, I, H, US, 20090101, 20090212, C, L)
    C07K-14/435(B, I, H, US, 20090101, 20090212, C, F)
    C07K-14/765(B, I, H, US, 20060101, 20090212, A, F)
    C08B-37/00(B,I,H,US,20060101,20090212,A,L)
    C08B-37/00(B, I, H, US, 20090101, 20090212, C, L)
Current US Class (main): 530-363000
Current US Class (secondary): 530-402000 536-124000
Original US Class (main): 530363
Original US Class (secondary): 536124 530402
Original Abstract: Three conjugation methods for use with the capsular
    saccharide of ~Streptococcus agalactiae~. In the first method,
    reductive animation of oxidised sialic acid residue side chains is
    used, but the aldehyde groups are first aminated, and then the amine is
    coupled to a carrier via a linker. In the second method, sialic acid
    residues and/or N-acetyl-glucosamine residues are de-N-acetylated to
    give amine groups, and the amine groups are coupled to a carrier
    protein via a linker. In the third method, linkage is via galactose
    residues in the capsular saccharide rather than sialic acid residues,
    which can conveniently be achieved using galactose oxidase.
Claim:
**1**. A process for preparing a conjugate of a ~Streptococcus
        agalactiae~ capsular saccharide and a carrier molecule, comprising
        the steps of: (a) oxidising a ~S. agalactiae~ capsular saccharide
        in order to introduce an aldehyde group into at least one terminal
        sialic acid residue in the saccharide; (b) subjecting the aldehyde
       group to reductive amination with ammonia or a primary amine, to
       give a --CH2-linked amine; (c) reacting the --CH2-linked amine with
       a bifunctional linker, to give an activated saccharide; and (d)
       reacting the activated saccharide with a carrier molecule, thereby
       giving the conjugate.
WIPO
Publication No. WO 2006082530 A2 (Update 200657 B)
Publication Date: 20060810
**CONJUGATION OF STREPTOCOCCAL CAPSULAR SACCHARIDES
  CONJUGAISON DE SACCHARIDES CAPSULAIRES STREPTOCOCCIOUES**
Assignee: ~(except US)~ CHIRON SRL, Via Fiorentina 1, I-53100 Siena, IT
    Residence: IT Nationality: IT (CHIR)
  ~(only US)~ BERTI, Francesco, Chiron Vaccines, Via Fiorentina, 1, I-53100
    Siena, IT Residence: IT Nationality: IT
Inventor: BERTI, Francesco, Chiron Vaccines, Via Fiorentina, 1, I-53100
    Siena, IT Residence: IT Nationality: IT
Agent: MARSHALL, Cameron, John et al., Carpmaels Ransford, 43-45
    Bloomsbury Square, London WC1A 2RA, GB
Language: EN (48 pages, 9 drawings)
Application: WO 2006IB756 A 20060201 (Local application)
Priority: GB 20052095 A 20050201
Designated States: (National Original) AE AG AL AM AT AU AZ BA BB BG BR BW
    BY BZ CA CH CN CO CR CU CZ DE DK DM DZ EC EE EG ES FI GB GD GE GH GM HR
    HU ID IL IN IS JP KE KG KM KN KP KR KZ LC LK LR LS LT LU LV LY MA MD MG
    MK MN MW MX MZ NA NG NI NO NZ OM PG PH PL PT RO RU SC SD SE SG SK SL SM
    SY TJ TM TN TR TT TZ UA UG US UZ VC VN YU ZA ZM ZW
  (Regional Original) AT BE BG BW CH CY CZ DE DK EA EE ES FI FR GB GH GM GR
    HU IE IS IT KE LS LT LU LV MC MW MZ NA NL OA PL PT RO SD SE SI SK SL SZ
    TR TZ UG ZM ZW
Original IPC: A61K-47/48(B,I,H,EP,20060101,A,F)
Current IPC: A61K-39/09(R,I,M,EP,20060101,20080531,A)
    A61K-39/09(R, I, M, EP, 20060101, 20080531, C)
   A61K-47/48(B, I, H, EP, 20060101, 20060810, A, F)
    A61K-47/48(B, I, H, EP, 20060101, 20060810, C, F)
Current ECLA class: A61K-39/09 A61K-47/48R2D A61K-47/48R2L A61K-47/48R2V
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Current ECLA ICO class: K61K-39:60P10

Original Abstract: Three conjugation methods for use with the capsular saccharide of Streptococcus agalactiae. In the first method, reductive animation of oxidised sialic acid residue side chains is used, but the aldehyde groups are first aminated, and then the amine is coupled to a carrier via a linker. In the second method, sialic acid residues and or N-acetyl-glucosamine residues are de-N-acetylated to give amine groups, and the amine groups are coupled to a carrier protein via a linker. In the third method, linkage is via galactose residues in the capsular saccharide rather than sialic acid residues, which can conveniently be achieved using galactose oxidase.

L'invention concerne trois procedes de conjugaison destines a etre utilises avec les saccharides capsulaires de Streptococcus agalactiae. Dans le premier procede, l'amination reductrice de chaines laterales de restes d'acide sialique oxyde est utilisee, mais les groupes aldehyde sont tout d'abord amines, puis l'amine est couplee a un support via un lieur. Dans le second procede, des residus d'acide sialique et/ou des residus de N-acetyl-glucosamine sont de-N-acetyl-les pour donner des groupes amine, et les groupes amine sont couples a une proteine via un lieur. Dans le troisieme procede, la liaison s'effectue via des residus de galactose dans le saccharide capsulaire plutot que dans les residus d'acide sialique qui peuvent etre facilement obtenus au moyen de galactose oxydase.

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galactose oxydase.
Publication No. WO 2006082530 A3 (Update 200845 E)
Publication Date: 20080703
Assignee: NOVARTIS VACCINES DIAGNOSTICS INC; IT (NOVS)
Inventor: BERTI F
Language: EN
Designated States: (National Original) AE AG AL AM AT AU AZ BA BB BG BR BW
    BY BZ CA CH CN CO CR CU CZ DE DK DM DZ EC EE EG ES FI GB GD GE GH GM HR
    HU ID IL IN IS JP KE KG KM KN KP KR KZ LC LK LR LS LT LU LV LY MA MD MG
    MK MN MW MX MZ NA NG NI NO NZ OM PG PH PL PT RO RU SC SD SE SG SK SL SM
    SY TJ TM TN TR TT TZ UA UG US UZ VC VN YU ZA ZM ZW
  (Regional Original) AT BE BG BW CH CY CZ DE DK EA EE ES FI FR GB GH GM GR
    HU IE IS IT KE LS LT LU LV MC MW MZ NA NL OA PL PT RO SD SE SI SK SL SZ
    TR TZ UG ZM ZW
Original IPC: A61K-39/09(B,I,H,EP,20060101,A,L)
    A61K-39/09(B,I,M,98,20060101,C) A61K-47/48(B,I,H,EP,20060101,A,F)
    A61K-47/48(B,I,M,98,20060101,C) A61P-37/00(B,I,M,98,20060101,C)
   A61P-37/04(B, I, H, EP, 20060101, A, L)
Current IPC: A61K-39/09(B,I,H,EP,20060101,20080703,A,L)
    A61K-39/09(B, I, H, EP, 20060101, 20080703, C, L)
    A61K-47/48(B, I, H, EP, 20060101, 20080703, A, F)
    A61K-47/48(B, I, H, EP, 20060101, 20080703, C, F)
    A61P-37/00(B,I,H,EP,20060101,20080703,C,L)
    A61P-37/04(B, I, H, EP, 20060101, 20080703, A, L)
Current ECLA class: A61K-39/09 A61K-47/48R2D A61K-47/48R2L A61K-47/48R2V
Current ECLA ICO class: K61K-39:60P10
South Africa
Publication No. ZA 200706969 A (Update 200918 E)
Publication Date: 20090128
Assignee: NOVARTIS VACCINES&DIAGNOSTICS INC (NOVS)
Inventor: BERTI F
Language: EN (56 pages)
Application: ZA 20076969 A 20070820 (Local application)
Priority: GB 20052095 A 20050201
Original IPC: A61K(A)
Current IPC: A61K(B,A,I,H,ZA,20090101,20090820,S)
   A61K-39/09(R,I,M,EP,20060101,20080531,A)
    A61K-39/09(R, I, M, EP, 20060101, 20080531, C)
    A61K-47/48(R, I, M, EP, 20060101, 20060722, A)
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A61K-47/48(R, I, M, EP, 20060101, 20060722, C) Current ECLA class: A61K-39/09 A61K-47/48R2D A61K-47/48R2L A61K-47/48R2V Current ECLA ICO class: K61K-39:60P10

10/7/13 (Item 13 from file: 351) DIALOG(R)File 351:Derwent WPI (c) 2011 Thomson Reuters. All rts. reserv.

0016018634 WPI ACC NO: 2006-550264/200656 XRAM Acc No: C2006-171980

Purifying Streptococcus agalactiae ****capsular**** ****polysaccharide**** involves treating suspension of streptococcal proteins, nucleic acids and ****polysaccharide**** with aqueous metal cation and alcohol, and treating aqueous material with cationic detergent

Patent Assignee: CHIRON SRL (CHIR); NOVARTIS VACCINES & DIAGNOSTICS INC (NOVS); NOVARTIS VACCINES&DIAGNOSTICS INC (NOVS); COSTANTINO P (COST-I) Inventor: COSTANTINO P; PAOLO C

Patent Family (11 patents, 112 countries)										
Patent Application										
Number Kind Date 1					Kind	Date	Update			
		2006082527	A2	20060810		2006IB626	A	20060201	200656	В
	EP	1848746	A2	20071031	EP	2006710574	A	20060201	200771	E
					WO	2006IB626	A	20060201		
	AU	2006211049	A1	20060810	AU	2006211049	A	20060201	200801	E
	CN	101146829	A	20080319	CN	200680007341	A	20060201	200841	E
					WO	2006IB626	A	20060201		
	JP	2008528052	W	20080731	WO	2006IB626	A	20060201	200853	E
					JP	2007553742	A	20060201		
	MX	2007009276	A1	20070901	WO	2006IB626	A	20060201	200864	E
					MX	20079276	A	20070801		
	za	200706968	A	20081126	ZA	20076968	A	20070820	200914	E
	US	20100063270	A1	20100311	WO	2006IB626	A	20060201	201019	E
					US	2008883615	A	20080512		
	NZ	560928	A	20100528	NZ	560928	A	20060201	201050	E
					WO	2006IB626	A	20060201		
	EP	2270056	A2	20110105	EP	2006710574	A	20060201	201104	E
					EP	2010179778	A	20060201		
	EP	2270056	A3	20110302	EP	2006710574	A	20060201	201117	E
					EP	2010179778	A	20060201		

Priority Applications (no., kind, date): GB 20052096 A 20050201

Patent Details Number Kind Lan Pg Dwg Filing Notes WO 2006082527 A2 EN 39 10 National Designated States, Original: AE AG AL AM AT AU AZ BA BB BG BR BW BY BZ CA CH CN CO CR CU CZ DE DK DM DZ EC EE EG ES FI GB GD GE GH GM HR HU ID IL IN IS JP KE KG KM KN KP KR KZ LC LK LR LS LT LU LV LY MA MD MG MK MN MW MX MZ NA NG NI NO NZ OM PG PH PL PT RO RU SC SD SE SG SK SL SM SY TJ TM TN TR TT TZ UA UG US UZ VC VN YU ZA ZM ZW Regional Designated States, Original: AT BE BG BW CH CY CZ DE DK EA EE ES FI FR GB GH GM GR HU IE IS IT KE LS LT LU LV MC MW MZ NA NL OA PL PT RO SD SE SI SK SL SZ TR TZ UG ZM ZW EP 1848746 A2 EN PCT Application WO 2006IB626

Based on OPI patent WO 2006082527 Regional Designated States, Original: AL AT BA BE BG CH CY CZ DE DK EE ES FI FR GB GR HR HU IE IS IT LI LT LU LV MC MK NL PL PT RO SE SI SK TR YU AU 2006211049 A1 EN Based on OPI patent WO 2006082527 CN 101146829 A ZH PCT Application WO 2006IB626 Based on OPI patent WO 2006082527

JP 2008528052 W JA 3.8 PCT Application WO 2006IB626 Based on OPI patent WO 2006082527 MX 2007009276 A1 ES PCT Application WO 2006IB626 Based on OPI patent WO 2006082527 ZA 200706968 EM 44 A US 20100063270 A1 EN PCT Application WO 2006IB626 NZ 560928 EN PCT Application WO 2006IB626 Based on OPI patent WO 2006082527 EP 2270056 A2 EN Division of application EP 2006710574

Division of patent EP 1848746

Regional Designated States,Original: AL AT BA BE BS CH CY CZ DE DK EE ES FI FR GB GR HR HU IE IS IT LI LT LU LV MC MK NL PL PT RO SE SI SK TR YU EP 2270056 A3 EN Division of application EP 2006710574

Division of patent EP 1848746

Regional Designated States, Original: AT BE BG CH CY CZ DE DK EE ES FI FR GB GR HU IE IS IT LI LT LU LV MC NL PL PT RO SE SI SK TR AL BA HR MK YU

Alerting Abstract WO A2

NOVELTY - Purification of ~Streptococcus agalactiae ~ ****capsular****
*****polysaccharide**** involves treating a suspension comprising
streptococcal proteins, nucleic acids and ****capsular****
*****polysaccharide**** with an aqueous metal cation and an alcohol to

****polysaccharide**** with an aqueous metal cation and an alcohol to precipitate nucleic acids and proteins; separating the precipitated material from the aqueous material; and treating the aqueous material with a cationic detergent to precipitate the ****capsular**** ****polysaccharide***.

DESCRIPTION - An INDEPENDENT CLAIM is also included for a composition comprising the purified ~Streptococcus agalactiae ~ ****capsular****

****polysaccharide****.

ACTIVITY - Antibacterial.

No biological data given.

MECHANISM OF ACTION - Vaccine.

USE - For purification of ~Streptococcus agalactiae ~ ****capsular****
****polysaccharide**** (claimed) useful in vaccine for bacterial
infections.

ADVANTAGE - The process avoids the need for DNase, RNase and/or protease treatment; is completed in less than three days after release of saccharide from the bacteria and has yield of about 60 (preferably 90)%. The saccharides have a very low protein contamination and a very low absorbance at 280 nm. The purity of the saccharide is at least 89 (preferably >=98)%.

Technology Focus

BIOTECHNOLOGY - Preferred Composition: The composition has UV absorbance at 280 nm of less than 0.20. The ratio of UV absorbance of the composition at 280 nm to the UV absorbance at 260 nm is greater than 0.85. The UV absorbance spectrum of the composition between 220 - 300 nm does exhibit either a shoulder or peak at around 270 nm. The UV spectrum of the composition between 250 - 275 nm has neither a maximum point nor a point of inflexion. The purity of the saccharide is at least 89% relative to the total weight of saccharide, protein and nucleic acid in the composition.

Preferred Components: The ****polysaccharide**** is from ~S. agalactiae ~ serotype selected from Ia, Ib, II, III, IV, V, VI, VII or VIII (preferably Ia, Ib, II, III or V). The ****polysaccharide**** is a full-length *****capsular*********polysaccharide**** and has a molecular weight greater than 30 kba. The saccharide is pertially or fully de-O-acetylated or de-N-acetylated. The suspension is the supernatant from a centrifuged ~S. agalactiae ~ culture and is prepared by treating (preferably chemically, enzymatically or by base extraction), ~S. agalactiae ~ such that the *****capsular**** saccharide is released. The *****capsular**** saccharide is released by treatment with both mutanolysin and

beta-N-acetylglucosaminidase, or by treatment with a type II phosphodiesterase.

INORGANIC CHEMISTRY - The aqueous metal cation is monovalent or divalent (preferably Mg ++ , Mn 4 ++ or Ca ++ (preferably Ca ++ (10 - 500 mM)). The aqueous medium comprises Mg ++ , Mn ++ or Ca ++ .

ORGANIC CHEMISTRY - Preferred Method: The alcohol is added to the suspension to give a final alcohol concentration of 10 - 50%. The precipitate is separated by centrifugation. The supernatant after centrifugation is subjected to microfiltration. A step of diafiltration is performed after step (a) and before step (c). The method further involves re-solubilizing the saccharide into aqueous medium or into alcoholic medium. The alcoholic medium has a final concentration of 70 - 95%.

Preferred Components: The alcohol is a lower alcohol (preferably ethanol or isopropanol). The cationic salt in step (c) is a tetrabutylammonium or cetyltrimethylammonium salt, such as cetyltrimethylammonium bromide.

Title Terms/Index Terms/Additional Words: PURIFICATION, STREFTOCCCUS; CAPSULE, ****POLYSACCHARIDE****; TREAT, SUSPENSION; PROTEIN; NUCLEIC;

ACID; AQUEOUS; METAL; CATION; ALCOHOL; MATERIAL; DETERGENT

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Class Codes
International Classification (+ Attributes)
IPC + Level Value Position Status Version
 A61K-0039/02 A I L B 20060101
  A61P-0031/04 A I L B 20060101
 A61F-0031/00 A I L B 20060101

C08B-0037/00 A I F 20060101

C08B-0037/00 A I F 20060101

C08B-0037/00 A I F B 20060101

C12P-0019/04 A I F B 20060101

C12P-0019/04 A I F B 20060101
 C12P-0019/04 A I L
                             20060101
 C12P-0019/04 A I
                          R 20060101
  C12P-0019/04 A I L B 20060101
 A61K-0039/02 C I L B 20060101
  A61P-0031/00 C I L B 20060101
 C08B S I B 20090101
 C08B-0037/00 C I L B 20060101
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  C08B-0037/00 C I
                          B 20060101
  C08B-0037/00 C I F B 20100101
  C12P S I B 20090101
  C12P-0019/00 C I F B 20060101
 C12P-0019/00 C I
 C12P-0019/00 C I
                          R 20060101
 C12P-0019/00 C I L B 20100101
ECLA: C08B-037/00K, C08B-037/00P, C12P-019/04
US Classification, Current Main: 536-123100
US Classification, Issued: 536123.1
JP Classification
  FI Term
                     Facet Rank Type
A61K-039/02
A61P-031/04
C08B-037/00
C12P-019/04
                    C ZNA
F-Term View Point Additional
 Theme
        + Figure Code
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 40085
 4C090
 4C201
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          CE06
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          CE11
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          DA01
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          DA 23
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File Segment: CPI
DWPI Class: B04; D16
Manual Codes (CPI/A-M): B04-C02F; B04-E01; B04-L05; B04-N03; B05-A01B;
 B05-A03A1; B10-A22; B10-E04D; B11-B03; B14-S11B1; D05-A02C; D05-H07;
 D05-H13
Original Publication Data by Authority
Australia
Publication No. AU 2006211049 A1 (Update 200801 E)
Publication Date: 20060810
Assignee: NOVARTIS VACCINES & DIAGNOSTICS INC (NOVS)
Inventor: COSTANTINO P
Language: EN
Application: AU 2006211049 A 20060201 (Local application)
Priority: GB 20052096 A 20050201
Related Publication: WO 2006082527 A (Based on OPI patent )
Original IPC: C08B-37/00(B,I,H,EP,20060101,20061116,A,F)
    C12P-19/04(B, I, H, EP, 20060101, 20061116, A, L)
Current IPC: C08B-37/00(B,I,H,EP,20060101,20061116,A,F)
    C08B-37/00(B,I,H,EP,20060101,20061116,C,F)
    C12P-19/00(B,I,H,EP,20060101,20061116,C,L)
    C12P-19/04(B, I, H, EP, 20060101, 20061116, A, L)
Current ECLA class: C08B-37/00K C08B-37/00P C12P-19/04
China
Publication No. CN 101146829 A (Update 200841 E)
Publication Date: 20080319
**Purification of streptococcal capsular polysaccharide**
Assignee: CHIRON SRL; IT (CHIR)
Inventor: PAOLO C
Language: ZH
Application: CN 200680007341 A 20060201 (Local application)
 WO 2006IB626 A 20060201 (PCT Application)
Priority: GB 20052096 A 20050201
Related Publication: WO 2006082527 A (Based on OPI patent )
Original IPC: C08B-37/00(I,CN,20060101,A,F) C08B-37/00(I,M,98,20060101,C)
    C12P-19/00(I,M,98,20060101,C) C12P-19/04(I,CN,20060101,A,L)
Current IPC: C08B-37/00(B,I,H,CN,20060101,20080319,A,F)
    C08B-37/00(B,I,H,CN,20060101,20080319,C,F)
    C12P-19/00(B, I, H, CN, 20060101, 20080319, C, L)
    C12P-19/04(B, I, H, CN, 20060101, 20080319, A, L)
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Current ECLA class: C08B-37/00K C08B-37/00P C12P-19/04

- Original Abstract: A purification method for the capsular polysaccharide of streptococcus agalactiae, wherein, the saccharide is initially treated by an aqueous mixture of an alcohol and a calcium salt, then it is processed with precipitation by a cationic detergent. This method can be completed in less than three days and has a yield of around 60%. It avoids the need for DNase, RNase and/or protease treatment. The saccharides obtained by this method have a very low protein contamination and a very low absorbance at 280nm.
- Claim: [CLAIM 1] A purification method for the capsular polysaccharide of streptococcus agalactiae, including following steps, (a) using an aqueous positive ion and alcohol to treat the mixture suspension containing streptococcus protein, nucleic acid and capsular polysaccharide to deposit nucleic acid and protein; (b) separating the deposited substance and aqueous substance; and (c) using a cationic detergent to treat the aqueous substance so as to deposit the capsular polysaccharide.
 - [CLAIM 2] The method according to claim 1, wherein, said polysaccharose is selected from none-streptococcus lactis serological type of Ia, Ib, II, III, IV, V, VI, VII or VIII.
 - [CLAIM 3] The method according to claim 2, wherein, said serological type is selected from Ia, Ib, II, III or V.
 - [CLAIM 4] The method according to any one of said claims, wherein, said polysaccharose is basically capsular polysaccharide with whole length. [CLAIM 5] The method according to any one of said claims, wherein,
 - molecular weight of said polysaccharose is more than 30 kDa.
 - [CLAIM 6] The method according to any one of said claims, wherein, said saccharides are partially or completely deoxy-acetylated.
 - [CLAIM 7] The method according to any one of said claims, wherein, said saccharides are partially or completely denitrifying-acetylated.
 - [CLAIM 8] The method according to any one of said claims, wherein, said mixture suspension is supernatant fluid of centrifugal streptococcus agalactiae culture.
 - [CLÄIM 9] The method according to any one of claims 1-7, wherein, the capsular polysaccharide is released by treating the streptococcus agalactiae so as to prepare said mixture suspension.
 - [CLAIM 10] The method according to claim 9, wherein, said capsular polysaccharide is released by a chemical treatment or an enzyme treatment.
 - [CLAIM 11] The method according to claim 10, wherein, said capsular polysaccharide is released by a alkali abstraction.
 - [CLAIM 12] The method according to claim 10, wherein, said capsular polysaccharide is released by a treatment using mutanolysin and beta-N-acetyl amidogen heteroside enzyme.
 - [CLAIM 13] The method according to claim 10, wherein, said capsular polysaccharide is released by a treatment of II type phosphodiesterase.
 - [CLAIM 14] The method according to any one of said claims, wherein, said alcohol is a low-grade alcohol.
 - [CLAIM 15] The method according to claim 14, wherein, said alcohol is ethanol or isopropanol.
 - [CLAIM 16] The method according to any one of said claims, wherein, said mixture suspension is added with said alcohol until concentration of the alcohol is between 10-50%.
 - [CLAIM 17] The method according to any one of said claims, wherein, said aqueous metal positive ion is monovalent or divalent.
 - [CLAIM 18] The method according to claim 17, wherein, said positive ion is Mg++, Mn++ or Ca++.
 - [CLAIM 19] The method according to claim 18, wherein, Ca++ is used, the finally concentration of it is between 10-500mM.
 - [CLAIM 20] The method according to any one of said claims, wherein, step (b) includes centrifugation.
 - [CLAIM 21] The method according to claim 20, wherein, the centrifugal

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supernatant fluid is filtered through a micro-hole.
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- [CLAIM 22] The method according to any one of said claims, wherein, a percolation step is performed after the step (a) and before the step
- [CLAIM 23] The method according to any one of said claims, wherein, the positive ion step in step (c) is tetrabutyl ammonium salt or hexadecyl trimethyl ammonium salt, such as CTAB.
- [CLAIM 24] The method according to any one of said claims, wherein, said method further includes re-dissolving said saccharides into aqueous medium or alcohol medium.
- [CLAIM 25] The method according to claim 24, wherein, using aqueous medium to re-dissolving said saccharides, wherein said aqueous medium contains Mg++, Mn++ or Ca++.
- [CLAIM 26] The method according to claim 24, wherein, using alcohol medium to re-dissolving said saccharides, wherein the final concentration of the alcohol is between 70-95%.
- [CLAIM 27] A combination containing streptococcus agalactiae obtained by the method according to any one of said claims.
- [CLAIM 28] The combination according to claim 27, wherein, 280 nm ultraviolet absorbency of said combination is less than 0.2.
- [CLAIM 29] The combination according to claim 27, wherein, a ratio between 280 nm ultraviolet absorbency of said combination and 260 nm ultraviolet absorbency us more than 0.85.
- [CLAIM 30] The combination according to claim 27, wherein, ultraviolet absorbency spectrum of said combination between 220-300 nm is about 270nm, and it shows an acromion or a peak.
- [CLAIM 31] The combination according to claim 27, wherein, ultraviolet absorbency spectrum of said combination between 250-275nm has no highest point or flex point.
- [CLAIM 32] The combination according to claim 27, wherein, relative to whole weight of the saccharides, protein and nucleic acid in the combination, purity quotient of said saccharides is at least 89%.

EPO

- Publication No. EP 1848746 A2 (Update 200771 E)
- Publication Date: 20071031
- **REINIGUNG VON STREPTOCOCCUS-KAPSELPOLYSACCHARID PURIFICATION OF STREPTOCOCCAL CAPSULAR POLYSACCHARIDE
- PURIFICATION DE POLYSACCHARIDES CAPSULAIRES STREPTOCOCCIQUES**
- Assignee: Novartis Vaccines and Diagnostics S.r.l., Via Fiorentina 1, 53100 Siena (SI), IT (NOVS)
- Inventor: COSTANTINO, PAOLO, CHIRON VACCINES, VIA FLORENTINA 1, I-53100 Siena, IT
- Agent: Marshall, Cameron John, Carpmaels Ransford, 43-45 Bloomsbury Square, London WC1A 2RA, GB

Language: EN

- Application: EP 2006710574 A 20060201 (Local application)
- WO 2006IB626 A 20060201 (PCT Application)
- Priority: GB 20052096 A 20050201
- Related Publication: WO 2006082527 A (Based on OPI patent)
- Designated States: (Regional Original) AL AT BA BE BG CH CY CZ DE DK EE ES FI FR GB GR HR HU IE IS IT LI LT LU LV MC MK NL PL PT RO SE SI SK TR YU
- Original IPC: C08B-37/00(B,I,H,EP,20060101,20060814,A,F)
 - C08B-37/00(B, I, M, 98, 20060101, 20060814, C)
 - C12P-19/00(B, I, M, 98, 20060101, 20060814, C)
- C12P-19/04(B, I, H, EP, 20060101, 20060814, A, L)
- Current IPC: C08B-37/00(B,I,H,EP,20060101,20060814,A,F)
 - C08B-37/00(B,I,H,EP,20060101,20060814,C,F)
 - C12P-19/00(B, I, H, EP, 20060101, 20060814, C, L)
- C12P-19/04(B, I, H, EP, 20060101, 20060814, A, L) Current ECLA class: C08B-37/00K C08B-37/00P C12P-19/04
- Original Abstract: A purification process for the capsular polysaccharide

of S.agalactiae in which the saccharide is initially treated with an aqueous mixture of an alcohol and a calcium salt, followed by precipitation with a cationic detergent. The process can be completed in less than three days and has a yield of around 60%. It avoids the need for DNase, RNase and/or protease treatment. The saccharides of the process have a very low protein contamination and a very low absorbance at 280nm. Publication No. EP 2270056 A2 (Update 201104 E) Publication Date: 20110105 **Reinigung von Streptococcus-kapselpolysaccharid Purification of streptococcal capsular polysaccharide Purification du polysaccharide capsulaire de streptococcus** Assignee: Novartis Vaccines and Diagnostics S.r.l., Via Fiorentina 1, 53100 Siena (SI), IT (NOVS) Inventor: Costantino, Paolo, CHIRON VACCINES, VIA FLORENTINA 1, I-53100 Siena, IT Agent: Marshall, Cameron John, Carpmaels Ransford, One Southampton Row, London, WC1B 5HA, GB Language: EN Application: EP 2010179778 A 20060201 (Local application) EP 2006710574 A 20060201 (Division of application) Priority: GB 20052096 A 20050201 Related Publication: EP 1848746 A (Division of patent) Designated States: (Regional Original) AL AT BA BE BG CH CY CZ DE DK EE ES FI FR GB GR HR HU IE IS IT LI LT LU LV MC MK NL PL PT RO SE SI SK TR YU Original IPC: C08B-37/00(B,I,H,EP,20060101,20101126,A,F) C12P-19/04(B, I, H, EP, 20060101, 20101126, A, L) Current IPC: C08B-37/00(B,I,H,EP,20060101,20101126,A,F) C12P-19/04(B, I, H, EP, 20060101, 20101126, A, L) Current ECLA class: C08B-37/00K C08B-37/00P C12P-19/04 Original Abstract: A purification process for the capsular polysaccharide of ~S.agalactiae~ in which the saccharide is initially treated with an aqueous mixture of an alcohol and a calcium salt, followed by precipitation with a cationic detergent. The process can be completed in less than three days and has a yield of around 60%. It avoids the need for DNase, RNase and/or protease treatment. The saccharides of the process have a very low protein contamination and a very low absorbance at 280nm. Claim: 1.A process for purifying a ~Streptococcus agalactiae~ capsular polysaccharide, comprising the step of removing contaminating nucleic acids and/or proteins by the use of precipitation. Publication No. EP 2270056 A3 (Update 201117 E) Publication Date: 20110302 **Purification of streptococcal capsular polysaccharide** Assignee: NOVARTIS VACCINESDIAGNOSTICS INC; IT (NOVS) Inventor: COSTANTINO P. IT Language: EN Application: EP 2010179778 A 20060201 (Local application) EP 2006710574 A 20060201 (Division of application) Priority: GB 20052096 A 20050201 Related Publication: EP 1848746 A (Division of patent)

C12P-19/04(B,I,H,EP,20060101,20101126,A,L)
Original Abstract: A purification process for the capsular polysaccharide of S.agalactiae in which the saccharide is initially treated with an

Designated States: (Regional Original) AT BE BG CH CY CZ DE DK EE ES FI FR GB GR HU IE IS IT LI LT LU LV MC NL PL PT RO SE SI SK TR AL BA HR MK YU

Original IPC: C08B-37/00(B,I,H,EP,20060101,20101126,A,F) C12P-19/04(B,I,H,EP,20060101,20101126,A,L) Current IPC: C08B-37/00(B,I,H,EP,20060101,20101126,A,F) aqueous mixture of an alcohol and a calcium salt, followed by precipitation with a cationic detergent. The process can be completed in less than three days and has a yield of around 60%. It avoids the need for DNase, RNase and/or protease treatment. The saccharides of the process have a very low protein contamination and a very low absorbance at 280mm.

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Publication No. JP 2008528052 W (Update 200853 E)
Publication Date: 20080731
Language: JA (38 pages)
Application: JP 2007553742 A 20060201 (Local application)
  WO 2006IB626 A 20060201 (PCT Application)
Priority: GB 20052096 A 20050201
Related Publication: WO 2006082527 A (Based on OPI patent )
Original IPC: A61K-39/02(B,I,H,JP,20060101,20080704,A,L)
   A61K-39/02(B, I, M, 98, 20060101, 20080704, C)
    A61P-31/00(B, I, M, 98, 20060101, 20080704, C)
    A61P-31/04(B, I, H, JP, 20060101, 20080704, A, L)
    C08B-37/00(B, I, H, JP, 20060101, 20080704, A, L)
    C08B-37/00(B, I, M, 98, 20060101, 20080704, C)
    C12P-19/00(B, I, M, 98, 20060101, 20080704, C)
    C12P-19/04(B, I, H, JP, 20060101, 20080704, A, F)
Current IPC: A61K-39/02(B.I.H.JP.20060101.20080704.A.L)
    A61K-39/02(B, I, H, JP, 20060101, 20080704, C, L)
    A61P-31/00(B, I, H, JP, 20060101, 20080704, C, L)
    A61P-31/04(B, I, H, JP, 20060101, 20080704, A, L)
    C08B-37/00(B, I, H, JP, 20060101, 20080704, A, L)
    C08B-37/00(B, I, H, JP, 20060101, 20080704, C, L)
    C12P-19/00(B, I, H, JP, 20060101, 20080704, C, F)
    C12P-19/04(B, I, H, JP, 20060101, 20080704, A, F)
Current ECLA class: C08B-37/00K C08B-37/00P C12P-19/04
Current JP F-Terms: 4B064 4C085 4C090 4C201 4C085AA03 4C090AA04 4C090AA09
    4B064AF11 4C085BA14 4C090BA94 4C090BC25 4C090BD37 4B064CA02 4C090CA18
    4C085CC07 4B064CC15 4B064CE03 4B064CE06 4B064CE11 4B064DA01 4C090DA23
    4C085DD37
Publication No. MX 2007009276 A1 (Update 200864 E)
Publication Date: 20070901
Assignee: NOVARTIS VACCINES & DIAGNOSTICS INC (NOVS)
Inventor: COSTANTINO P
Language: ES
Application: WO 2006IB626 A 20060201 (PCT Application)
 MX 20079276 A 20070801 (Local application)
Priority: GB 20052096 A 20050201
Related Publication: WO 2006082527 A (Based on OPI patent )
Original IPC: C08B-37/00(I,MX,20060101,A,F) C08B-37/00(I,M,98,20060101,C)
    C12P-19/00(I,M,98,20060101,C) C12P-19/04(I,MX,20060101,A,L)
Current IPC: C08B-37/00(I,MX,20060101,A,F) C08B-37/00(I,M,98,20060101,C)
    C12P-19/00(I,M,98,20060101,C) C12P-19/04(I,MX,20060101,A,L)
New Zealand
Publication No. NZ 560928 A (Update 201050 E)
Publication Date: 20100528
Assignee: NOVARTIS VACCINES & DIAGNOSTICS INC (NOVS)
Inventor: COSTANTINO P
Language: EN
Application: NZ 560928 A 20060201 (Local application)
 WO 2006IB626 A 20060201 (PCT Application)
Priority: GB 20052096 A 20050201
Related Publication: WO 2006082527 A (Based on OPI patent )
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Original IPC: C08B-37/00(I,NZ,20060101,A,F) C08B-37/00(I,M,98,20060101,C)
    C12P-19/00(I,M,98,20060101,C) C12P-19/04(I,NZ,20060101,A,L)
Current IPC: C08B-37/00(B,I,H,NZ,20060101,20100622,A,F)
    C08B-37/00(B, I, H, NZ, 20100101, 20100622, C, F)
    C12P-19/00(B, I, H, NZ, 20100101, 20100622, C, L)
    C12P-19/04(B, I, H, NZ, 20060101, 20100622, A, L)
Current ECLA class: C08B-37/00K C08B-37/00P C12P-19/04
United States
Publication No. US 20100063270 A1 (Update 201019 E)
Publication Date: 20100311
**Purification of Streptococcal Capsular Polysaccharide**
Assignee: Costantino, Paolo, Siena, IT Residence: IT (COST-I)
Inventor: Costantino, Paolo, Siena, IT Residence: IT
Agent: NOVARTIS VACCINES AND DIAGNOSTICS INC., INTELLECTUAL PROPERTY-
    X100B, P.O. BOX 8097, Emeryville, CA, US
Language: EN
Application: US 2008883615 A 20080512 (Local application)
  WO 2006IB626 A 20060201 (PCT Application)
Priority: GB 20052096 A 20050201
Original IPC: C08B-37/00(B,I,H,US,20060101,20100311,A,F)
    C08B-37/00(B, I, M, 98, 20060101, 20100311, C)
Current IPC: C08B-37/00(B,I,H,US,20060101,20100311,A,F)
    C08B-37/00(B, I, M, 98, 20060101, 20100311, C)
Current ECLA class: C08B-37/00K C08B-37/00P C12P-19/04
Current US Class (main): 536-123100
Original US Class (main): 536123.1
Original Abstract: A purification process for the capsular polysaccharide
    of ~S. agalactiae~ in which the saccharide is initially treated with an
    aqueous mixture of an alcohol and a calcium salt, followed by
    precipitation with a cationic detergent. The process can be completed
    in less than three days and has a yield of around 60%. It avoids the
    need for DNase, RNase and/or protease treatment. The saccharides of the
    process have a very low protein contamination and a very low absorbance
    at 280 nm.
Claim:
**1**. A process for purifying a ~Streptococcus agalactiae~ capsular
        polysaccharide, comprising the steps of: (a) treating a suspension
        comprising streptococcal proteins, nucleic acids and capsular
        polysaccharide with an aqueous metal cation and an alcohol in order
        to precipitate nucleic acids and proteins; (b) separating the
        precipitated material from the aqueous material; and (c) treating
        the aqueous material with a cationic detergent in order to
        precipitate the capsular polysaccharide.
Publication No. WO 2006082527 A2 (Update 200656 B)
Publication Date: 20060810
**PURIFICATION OF STREPTOCOCCAL CAPSULAR POLYSACCHARIDE
  PURIFICATION DE POLYSACCHARIDES CAPSULAIRES STREPTOCOCCIQUES**
Assignee: ~(except US)~ CHIRON SRL, Via Fiorentina 1, I-53100 Siena, IT
    Residence: IT Nationality: IT (CHIR)
  ~(only US)~ COSTANTINO, Paolo, Chiron Vaccines, Via Florentina 1, I-53100 Siena, IT Residence: IT Nationality: IT
Inventor: COSTANTINO, Paolo, Chiron Vaccines, Via Florentina 1, I-53100
    Siena, IT Residence: IT Nationality: IT
Agent: MARSHALL, Cameron, John et al., Carpmaels Ransford, 43-45
    Bloomsbury Square, London WC1A 2RA, GB
Language: EN (39 pages, 10 drawings)
Application: WO 2006IB626 A 20060201 (Local application)
Priority: GB 20052096 A 20050201
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Designated States: (National Original) AE AG AL AM AT AU AZ BA BB BG BR BW
    BY BZ CA CH CN CO CR CU CZ DE DK DM DZ EC EE EG ES FI GB GD GE GH GM HR
    HU ID IL IN IS JP KE KG KM KN KP KR KZ LC LK LR LS LT LU LV LY MA MD MG
    MK MN MW MX MZ NA NG NI NO NZ OM PG PH PL PT RO RU SC SD SE SG SK SL SM
    SY TJ TM TN TR TT TZ UA UG US UZ VC VN YU ZA ZM ZW
  (Regional Original) AT BE BG BW CH CY CZ DE DK EA EE ES FI FR GB GH GM GR
    HU IE IS IT KE LS LT LU LV MC MW MZ NA NL OA PL PT RO SD SE SI SK SL SZ
    TR TZ UG ZM ZW
Original IPC: C08B-37/00(B,I,H,EP,20060101,A,F)
    C12P-19/00(B,I,H,98,20060101,C,L) C12P-19/04(B,I,H,EP,20060101,A,L)
Current IPC: C08B-37/00(B,I,H,EP,20060101,20060810,A,F)
    C08B-37/00(B, I, H, EP, 20060101, 20060810, C, F)
    C12P-19/00(B, I, H, EP, 20060101, 20060810, C, L)
    C12P-19/04(B, I, H, EP, 20060101, 20060810, A, L)
Current ECLA class: C08B-37/00K C08B-37/00P C12P-19/04
Original Abstract: A purification process for the capsular polysaccharide
    of S.agalactiae in which the saccharide is initially treated with an
    aqueous mixture of an alcohol and a calcium salt, followed by
    precipitation with a cationic detergent. The process can be completed
    in less than three days and has a yield of around 60%. It avoids the
    need for DNase, RNase and/or protease treatment. The saccharides of the
    process have a very low protein contamination and a very low absorbance
    at 280nm.
  L'invention concerne un procede de purification pour polysaccharides
    capsulaires de type S. agalactiae, selon lequel les saccharides sont
    tout d'abord traites avec un melange aqueux a base d'alcool et de sel
    de calcium, ledit traitement etant suivi par la precipitation d'un
    detergent cationique. Le procede peut etre effectue en moins de trois
    jours et presente un rendement d'environ 60%. Ledit procede evite
    d'avoir recours a un traitement par DNase, RNase et ou protease. Les
   saccharides selon le procede presentent un taux tres reduit de
   contamination proteinique et un taux d'absorbance tres reduit a 280 nm.
South Africa
Publication No. ZA 200706968 A (Update 200914 E)
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Publication No. ZA 200706968 A (Update 200914 E)
Publication Date: 20081126
Assignee: NOVARTIS VACCINES&DIAGNOSTICS INC (NOVS)
Inventor: COSIANTINO PER PROPERTION (NOVS)
Language: EN (44 pages)
Application: ZA 20076968 A 20070820 (Local application)
Priority: GB 20052096 A 20050201
Original IPC: CO8B(A) Cl2P(B)
Current IPC: CO8B(A), I, H, ZA, 20090101, 20090812, S)
C08B-37/00(R, I, M, EP, 20060101, 20070721, A)
C08B-37/00(R, I, M, EP, 20060101, 20070721, C)
C12P(B, I, H, ZA, 20090101, 20090812, S)
C12P-19/00(R, I, M, EP, 20060101, 20070721, C)
C12P-19/00(R, I, M, EP, 20060101, 20070721, A)
Current ECLA class: C08B-37/00K C08B-37/00P C12P-19/04
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10/7/14 (Item 14 from file: 351)
DIALOG(R)File 351:Derwent WPI
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0014974548 - Drawing available
WPI ACC NO: 2005-322381/200533
XRAM Acc No: C2005-100521
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Modified serogroup meningococcal *****capsular**** saccharide useful for the preparation of immunogenic compositions have altered levels of ortho-acetylation at the specified positions of their sialic acid residues

Patent Assignee: CHIRON SRL (CHIR); NOVARTIS VACCINES&DIAGNOSTICS INC (NOVS); NOVARTIS VACCINES & DIAGNOSTICS INC (NOVS); BERTI F (BERT-I); COSTANTINO P (COST-I)

Inventor: COSTANTINO P: BERTI F

Patent Family (14 patents, 107 countries) Pat.ent. Application Number Kind Date Kind Date Number Updat.e A 20041004 200533 B WO 2005033148 A1 20050414 WO 2004IB3366 A1 20060712 EP 2004791697 EP 1678212 A 20041004 200648 E WO 2004IB3366 A 20041004 MX 2006003729 Al 20060701 WO 2004IB3366 A 20041004 200677 E A 20060403 MX 20063729 AU 2004278170 A1 20050414 AU 2004278170 A 20041004 200681 E BR 200415048 A 20061212 BR 200415048 A 20041004 200701 E W0 2004183366 A 20041004 200702 E U 20070507578 W 20070329 W0 2004183366 A 20041004 200725 E JP 2006530760 A 20041004 CN 1882612 A 20061220 CN 200480034525 A 20041004 200730 B
A1 20080710 AU 2004278170 A 20041004 200864 NCE
AU 2008202708 A 20080619
B2 20080703 AU 2004278170 A 20041004 200867 E
A 20090626 NZ 546669 A 20041004 200946 E
W0 20041B3366 A 20041004 200950 E
A1 20100415 W0 20041B3366 A 20041004 200950 E
A1 20100415 W0 20041B3366 A 20041004 201027 E
US 2007574437 A 20070425
B2 2010527 AU 2004278170 A 20041004 201039 NCE
AU 20011229 EP 2004791697 A 20041004 201104 E A 20061220 CN 200480034525 A 20041004 200730 E AU 2008202708 AU 2004278170 NZ 546668 RU 2362784 US 20100092509 AU 2008202708 EP 2267036 A1 20101229 EP 2004791697 A 20041004 201104 E EP 2010180746 A 20041004

Priority Applications (no., kind, date): GB 200323103 A 20031002; AU 2008202708 A 20080619

Patent Details

Number

Kind Lan Pg Dwg Filing Notes

WO 2005033148 A1 EN 42 6

National Designated States, Original: AE AG AL AM AT AU AZ BA BB BG BR BW BY BZ CA CH CN CO CR CU CZ DE DK DM DZ EC EE EG ES FI GB GD GE GH GM HR HU ID IL IN IS JP KE KG KP KR KZ LC LK LR LS LT LU LV MA MD MG MK MN MW MX MZ NA NI NO NZ OM PG PH PL PT RO RU SC SD SE SG SK SL SY TJ TM TN TR TT TZ UA UG US UZ VC VN YU ZA ZM ZW

Regional Designated States, Original: AT BE BG BW CH CY CZ DE DK EA EE ES FI FR GB GH GM GR HU IE IT KE LS LU MC MW MZ NA NL OA PL PT RO SD SE SI SK SL SZ TR TZ UG ZM ZW

EP 1678212 A1 EN

Regional Designated States, Original: AT BE BG CH CY CZ DE DK EE ES FI FR IT LI LU MC NL PL PT RO SE SI SK TR

MX	GB GR HU IE 2006003729		ES	MC	NL
	2004278170 200415048	A1 A	EN PT		
JP	2007507578	W	JA		42
ΑÜ	2008202708 2004278170 546668		EN EN EN		
RU	2362784	C2	RU		

PCT Application WO 2004IB3366 Based on OPI patent WO 2005033148

PCT Application WO 2004IB3366 Based on OPI patent WO 2005033148 WO 2005033148 Based on OPI patent PCT Application WO 2004IB3366 Based on OPI patent WO 2005033148 PCT Application WO 2004IB3366 Based on OPI patent WO 2005033148 Division of application AU 2004278170 Based on OPI patent WO 2005033148 PCT Application WO 2004IB3366 Based on OPI patent WO 2005033148 PCT Application WO 2004IB3366

Based on OPI patent W0 2005033148 US 20100092509 A1 EN PCT Application W0 2004IB3366 AU 2008202708 B2 EN Division of application AU 2004278170

EP 2267036 A1 EN Division of application EP 2004791697

Division of patent EP 1678212

Regional Designated States, Original: AT BE BG CH CY CZ DE DK EE ES FI FR GB GR HU IE IT LI LU MC NL PL PT RO SE SI SK TR

Alerting Abstract WO Al

NOVELTY - A modified serogroup W135 meningococcal *****capsular**** saccharide (I), where (a) <= 29% of the sialic acid residues in the saccharide are ortho-acetylated at the 7 position and (b)>= 26% of the sialic acid residues in the saccharide are ortho-acetylated at the 9 position.

DESCRIPTION - INDEPENDENT CLAIMS are included for the following:
1.a modified serogroup Y meningococcal ****capsular**** saccharide (II),
where <= 9% of the sialic acid residues in the saccharide are
Ortho-acetylated at the 7 position and/or (b) >= 29% or <= 27% of the
sialic acid residues in the saccharide are ortho-acetylated at the 9
position;

- 2.a modified meningococcal ****capsular**** saccharide (III), optionally conjugated to a carrier protein, where the saccharide comprises n or more repeating units of the disaccharide unit: [sialic acid]-[hexose];
- 3.a composition (C1) comprising (a1) molecules of serogroup W135 meningococal ***-capsular**** saccharide, where the average number of sialic acid residues per ****-capsular**** saccharide molecule is (b1), and where <= 29% of the (alxb1) serogroup W135 sialic acid residues in the composition are ortho-acetylated at the 7 position and/or 26% of the axb serogroup W135 sialic acid residues in the composition are ortho-acetylated at the 9 position;</p>
- 4.a composition (C2) comprising (a2) molecules of serogroup Y meningococal ****capsular**** saccharide, where the average number of sialic acid residues per ****capsular**** accharide molecule is (b2), and where the average number of sialic acid residues per ****capsular**** saccharide molecule is (b2) and where <= 9% of the a2xb2 serogroup Y sialic acid residues in the composition are ortho-acetylated at the 7 position and for >= 29% of <= 27% of the (a2)x(b2) serogroup Y sialic acid residues in the composition are ortho-acetylated at the 9 position;</p>
- 5.a saccharide (S1) comprising n or more repeats of the disaccharide unit of formula (Ia);
- 6.a conjugation product of (S1) and a carrier protein selected from diphtheria toxoid, tetanus toxoid, H. influenzae protein D, and CRM197;
- 7.an immunogenic composition (III) comprising the modified ****capsular**** saccharides or conjugates and a carrier; and
- 8.preparation of an immunogenic conjugate involving providing a starting serogroup W135 or serogroup Y meningococcal ****capsular**** saccharide and a carrier protein, either or both of which is/are optionally modified to render it/them reactive towards the other, forming a covalent bond between the saccharide and the carrier protein and purifying the resulting glycoconjugates, where, between the first and

the third step, the degree of ortho-acetylation at the 9 position of sialic acid residues in the starting saccharide increases.

<img

src="http://imagesrv.dialog.com/imanager/getimage?ref=Iaccde890b58e11daaa26000083613
46f&f=351&type=PNG" width="2513" height="2206"/> hexose= galactose or glucose;

n= 1 - 100;

X,Y= H or OH;

R1, R2 = H or -COCH 3.

Provided that:

- 1.(a) >= x% of the sialic acid residues in the n or more repeating units are ortho-acetylated at the 7 position and/or (b) when hexose is galactose, greater than y% of the sialic acid residues in the n or more repeating units are ortho-acetylated at the 9 position, and when hexose is glucose, >= y% or >= z% of the sialic acid residues in the n or more repeating units are ortho-acetylated at the 9 position, where when hexose is galactose, x is 29 and y is 26, and when hexose is glucose, x is 39, y is 29 and z is 27 (preferably when hexose is galactose, 6% of the sialic acid residues in the n or more repeating units are ortho-acetylated at the 7 position, and 43% of the sialic acid residues in the n or more repeating units are ortho-acetylated at the 9 position; and when hexose is glucose, 6% of the sialic acid residues in the n or more repeating units are ortho-acetylated at the 7 position, and 45% of the sialic acid residues in the n or more repeating units are ortho-acetylated at the 7 position, and 45% of the sialic acid residues in the n or more repeating units are ortho-acetylated at the 9 position); and
- 2.when X is OH and Y is H, <=29% of R 1 are COCH 3 and/or >= 26% of R 2 are COCH3; and when X is H and Y is OH, <= 9% of R 1 are COCH 3 and/or >= 29% of < 27% of R 2 are COCH3.

ACTIVITY - Antibacterial.

MECHANISM OF ACTION - None given.

USE - In the preparation of an immunogenic composition useful as a medicament for raising an antibody response; as a medicament for protecting against meningococcal meningitis (all claimed).

ADVANTAGE - The conjugates show improved immunogenicity compared to native polysaccharides.

Technology Focus

ORGANIC ČHEMISTRY — Preferred Composition: In (I) and (II), greater than 05 fthe sialic acid residues in the saccharide are ortho-acetylated at the 7 and 9 position. In (C1) and (C2), the***capsular**** saccharide is conjugated to a protein carrier. In (S1), the saccharide has an average degree of polymerization of less than 30. (III) is in aqueous form or in lyophilized form.

BIOLOGY - Preferred Composition: (III) further comprises a

****capsular**** saccharide antigen from serogroup C or A of ~N.

****meningitidis**** ~ , an antigen from serogroup B of ~N.

****meningitidis**** ~ , a saccharide antigen from ~Haemophilus influenzae type B ~ , an antigen from ~Streptococcus pneumoniae ~ , an antigen from hepatitis A virus, an antigen from hepatitis B virus, an antigen from ~Bordetella pertussis ~ , a diphtheria toxoid, a tetanus toxoid and/or a poliovirus antigen. The serogroup A is an antigen.

Title Terms/Index Terms/Additional Words: MODIFIED; MENINGCOCCUS; CAPSULE; SACCHARIDE; USEFUL; PREPARATION; IMMUNOGENIC; COMPOSITION; ALTER; LEVEL; ORTHO; ACETYLATE; SPECIFIED; POSITION; SIALIC; ACID; RESIDUE

Class Codes International Classification (Main): C08B-037/00 (Additional/Secondary): A61K-031/715, A61K-039/095 International Classification (+ Attributes) IPC + Level Value Position Status Version A61K-0031/715 A I F B 20060101 A61K-0031/715 A I B 20060101 A61K-0031/715 A I R 20060101 A61K-0039/05 A I L B 20060101 A61K-0039/08 A I L B 20060101 A61K-0039/09 A I L B 20060101 A61K-0039/095 A I F B 20060101 B 20060101 A61K-0039/095 A I A61K-0039/095 A I R 20060101 A61K-0039/10 A I L B 20060101

A61K-0039/102 A I L B 20060101 A61K-0039/16 A I L B 20060101 A61K-0039/13 A I L B 20060101 A61K-0039/29 A I L B 20060101 A61K-0039/29 A I L B 20060101 A61K-0039/385 A I F B 20060101 A61F-0001/16 A I L B 20060101 A61F-0031/04 A I L B 20060101 A61F-0031/4 A I L B 20060101 A61F-0031/4 A I L B 20060101 A61F-0031/20 A I L B 20060101 A61F-0031/20 A I L B 20060101 C07H-0001/00 A I L B 20060101 C07K-0014/22 A I L B 20060101 C07K-0014/22 A I L B 20060101

C07K-0014/22 A N L B 20060101 C08B-0037/00 A I L B 20060101 C08B-0037/00 A I B 20060101 C08B-0037/00 A I B 20060101 C12N-0015/09 A N L B 20060101

A61K-0039/05 C I L B 20060101 A61K-0039/08 C I L B 20060101 A61K-0039/09 C I L B 20060101 A61K-0039/09 C I L B 20060101 A61K-0039/100 C I L B 20060101 A61K-0039/102 C I L B 20060101

A61K-0039/116 C I L B 20100101 A61K-0039/29 C I L B 20060101 A61K-0039/29 C I L B 20060101 A61K-0039/385 C I F B 20100101 A61P-0001/00 C I L B 20060101

AGIP-0037/00 C I L B 20060101 C07H-0001/00 C I L B 20100101 C07K-0014/195 C I L B 20100101 C07K-0014/195 C N L B 20060101 C08B-0037/00 C I F B 20060101 C08B-0037/00 C I F B 20060101

C12N-0015/09 C N L B 20060101 A61K-0031/715 C I F B 20100101 A61K-0031/715 C I B 20060101

A61K-0039/095 C I L B 20100101 A61K-0039/095 C I B 20060101

A61P-0031/00 C I L B 20100101 C08B-0037/00 C I L B 20100101

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C08B-0037/00 C I B 20060101
ECLA: A61K-031/715, A61K-039/095, C08B-037/00P
ICO: K61K-039:60P10
US Classification, Current Main: 424-197110; Secondary: 424-203100,
424-250100, 530-395000, 536-123100
US Classification, Issued: 424197.11, 536123.1, 530395, 424250.1, 424203.1
JP Classification
  FI Term
                  Facet Rank Type
A61K-031/715
A61K-039/05
A61K-039/08
A61K-039/09
A61K-039/095
A61K-039/10
A61K-039/102
A61K-039/13
A61K-039/29
A61P-001/16
A61P-031/04
A61P-031/14
A61P-031/20
A61P-037/04
C07K-014/22
C08B-037/00
C08B-037/00
                   P
                     ZNA
C12N-015/00
                  Α
F-Term View Point Additional
 Theme
       + Figure Code
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          AA01
 4C086
          AA02
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          AA03
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          AA04
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          AA05
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          AA09
 4H045
          AA11
 4H045
          AA20
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          AA30
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          BA10
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          BA17
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4C090

4C085

BA94

BB11

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          CA11
           CA39
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 4C090
           CA46
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          CC07
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           CC08
 4B024
          DA05
 4C090
           DA09
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           DA 23
 4H045
          DA86
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          EA04
 4H045
          EA22
 4C086
          EA25
 4H045
          EA29
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           GG01
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           MA02
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          ZB35
File Segment: CPI
DWPI Class: A11; A96; B03; B04
Manual Codes (CPI/A-M): A03-A01; A10-E01; A12-V01; B04-C02; B14-A01A5;
  B14-G01
Original Publication Data by Authority
Australia
Publication No. AU 2004278170 A1 (Update 200681 E)
Publication Date: 20050414
Assignee: CHIRON SRL (CHIR)
Inventor: COSTANTINO P
Language: EN
Application: AU 2004278170 A 20041004 (Local application)
Priority: GB 200323103 A 20031002
Related Publication: WO 2005033148 A (Based on OPI patent )
Original IPC: A61K-31/715(B,I,M,EP,20060101,20060408,A,L)
    A61K-39/095(B, I, M, EP, 20060101, 20060408, A, F)
    C08B-37/00(B,I,M,EP,20060101,20060408,A,L)
Current IPC: A61K-31/715(B,I,M,EP,20060101,20060408,A,L)
    A61K-39/095(B, I, M, EP, 20060101, 20060408, A, F)
    C08B-37/00(B, I, M, EP, 20060101, 20060408, A, L)
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Current ECLA class: A61K-31/715 A61K-39/095 C08B-37/00P
Current ECLA ICO class: K61K-39:60P10
Publication No. AU 2004278170 B2 (Update 200867 E)
Publication Date: 20080703
Assignee: NOVARTIS VACCINES & DIAGNOSTICS INC (NOVS)
Inventor: COSTANTINO P
Language: EN
Application: AU 2004278170 A 20041004 (Local application)
Priority: GB 200323103 A 20031002
Related Publication: WO 2005033148 A (Based on OPI patent )
Original IPC: A61K-31/715(B,I,M,EP,20060101,20051008,A,F)
    A61K-31/715(B, I, M, 98, 20060101, 20051008, C)
    A61K-39/095(B, I, M, EP, 20060101, 20051008, A, L)
    A61K-39/095(B, I, M, 98, 20060101, 20051008, C)
    C08B-37/00(B, I, M, EP, 20060101, 20051008, A, L)
    C08B-37/00(B, I, M, 98, 20060101, 20051008, C)
Current IPC: A61K-31/715(B,I,M,EP,20060101,20051008,A,F)
    A61K-31/715(B, I, M, EP, 20060101, 20051008, C, F)
    A61K-39/095(B, I, M, EP, 20060101, 20051008, A, L)
    A61K-39/095(B, I, M, EP, 20060101, 20051008, C, L)
    C08B-37/00(B,I,M,EP,20060101,20051008,A,L)
    C08B-37/00(B,I,M,EP,20060101,20051008,C,L)
Current ECLA class: A61K-31/715 A61K-39/095 C08B-37/00P
Current ECLA ICO class: K61K-39:60P10
Publication No. AU 2008202708 Al (Update 200864 NCE)
Publication Date: 20080710
Assignee: NOVARTIS VACCINES & DIAGNOSTICS INC (NOVS)
Inventor: COSTANTINO P
Language: EN
Application: AU 2004278170 A 20041004 (Division of application)
  AU 2008202708 A 20080619 (Local application)
Priority: AU 2008202708 A 20080619 (Local application)
Original IPC: A61K-31/715(B,I,M,AU,20060101,20080624,A,F)
    A61K-31/715(B, I, M, 98, 20060101, 20080624, C)
    A61K-39/095(B, I, M, AU, 20060101, 20080624, A, L)
    A61K-39/095(B, I, M, 98, 20060101, 20080624, C)
    C08B-37/00(B, I, M, AU, 20060101, 20080624, A, L)
    C08B-37/00(B,I,M,98,20060101,20080624,C)
Current IPC: A61K-31/715(B,I,M,AU,20060101,20080624,A,F)
   A61K-31/715(B, I, M, AU, 20060101, 20080624, C, F)
    A61K-39/095(B, I, M, AU, 20060101, 20080624, A, L)
    A61K-39/095(B, I, M, AU, 20060101, 20080624, C, L)
    C08B-37/00(B, I, M, AU, 20060101, 20080624, A, L)
    C08B-37/00(B,I,M,AU,20060101,20080624,C,L)
Current ECLA class: A61K-31/715 A61K-39/095 C08B-37/00P
Current ECLA ICO class: K61K-39:60P10
Publication No. AU 2008202708 B2 (Update 201039 NCE)
Publication Date: 20100527
Assignee: NOVARTIS VACCINES & DIAGNOSTICS INC (NOVS)
Inventor: BERTI F
  COSTANTINO P
Language: EN
Application: AU 2008202708 A 20080619 (Local application)
  AU 2004278170 A 20041004 (Division of application)
Priority: AU 2008202708 A 20080619 (Local application)
Original IPC: A61K-31/715(B,I,M,AU,20060101,20080624,A,F)
    A61K-31/715(B,I,M,98,20060101,20080624,C)
    A61K-39/095(B, I, M, AU, 20060101, 20080624, A, L)
   A61K-39/095(B, I, M, 98, 20060101, 20080624, C)
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C08B-37/00(B, I, M, AU, 20060101, 20080624, A, L)
    C08B-37/00(B, I, M, 98, 20060101, 20080624, C)
Current IPC: A61K-31/715(B,I,M,AU,20060101,20080624,A,F)
    A61K-31/715(B, I, M, AU, 20100101, 20080624, C, F)
    A61K-39/095(B, I, M, AU, 20060101, 20080624, A, L)
    A61K-39/095 (B, I, M, AU, 20100101, 20080624, C, L)
    C08B-37/00(B, I, M, AU, 20060101, 20080624, A, L)
    C08B-37/00(B, I, M, AU, 20100101, 20080624, C, L)
Current ECLA class: A61K-31/715 A61K-39/095 C08B-37/00P
Current ECLA ICO class: K61K-39:60P10
Publication No. BR 200415048 A (Update 200701 E)
Publication Date: 20061212
Assignee: CHIRON SRL (CHIR)
Inventor: COSTANTINO P
Language: PT
Application: BR 200415048 A 20041004 (Local application)
  WO 2004IB3366 A 20041004 (PCT Application)
Priority: GB 200323103 A 20031002
Related Publication: WO 2005033148 A (Based on OPI patent )
Original IPC: C08B-37/00(A) A61K-31/715(B) A61K-39/095(B)
Current IPC: C08B-37/00(A) A61K-31/715(B) A61K-39/095(B)
Current ECLA class: A61K-31/715 A61K-39/095 C08B-37/00P
Current ECLA ICO class: K61K-39:60P10
China
Publication No. CN 1882612 A (Update 200730 E)
Publication Date: 20061220
Language: ZH
Application: CN 200480034525 A 20041004 (Local application)
Priority: GB 200323103 A 20031002
Original IPC: A61K-31/715(I,CN,20060101,A,L) A61K-31/715(I,M,98,20060101,C)
    A61K-39/095(I,CN,20060101,A,L) A61K-39/095(I,M,98,20060101,C)
    C08B-37/00(I,CN,20060101,A,F) C08B-37/00(I,M,98,20060101,C)
Current IPC: A61K-31/715(B, I, H, CN, 20060101, 20061220, A, L)
    A61K-31/715(B, I, H, CN, 20060101, 20061220, C, L)
    A61K-39/095(B, I, H, CN, 20060101, 20061220, A, L)
    A61K-39/095(B, I, H, CN, 20060101, 20061220, C, L)
    C08B-37/00(B, I, H, CN, 20060101, 20061220, A, F)
    C08B-37/00(B,I,H,CN,20060101,20061220,C,F)
Current ECLA class: A61K-31/715 A61K-39/095 C08B-37/00P
Current ECLA ICO class: K61K-39:60P10
EPO
Publication No. EP 1678212 Al (Update 200648 E)
Publication Date: 20060712
**HYPO- UND HYPERACETYLIERTE MENINGOKOKKEN-KAPSELSACCHARIDE
 HYPO- AND HYPER-ACETYLATED MENINGOCOCCAL CAPSULAR SACCHARIDES
  SACCHARIDES CAPSULAIRES MENINGOCOCCIQUES HYPO ET HYPERACETYLES**
Assignee: Chiron SRL., Via Fiorentina, 1, 53100 Siena, IT
Inventor: COSTANTINO, Paolo, Chiron SRL, Via Fiorentina 1, I-53100 Siena,
Agent: Marshall, Cameron John, Carpmaels Ransford, 43-45 Bloomsbury
    Square, London WC1A 2RA, GB
Language: EN
Application: EP 2004791697 A 20041004 (Local application)
  WO 2004IB3366 A 20041004 (PCT Application)
Priority: GB 200323103 A 20031002
Related Publication: WO 2005033148 A (Based on OPI patent )
Designated States: (Regional Original) AT BE BG CH CY CZ DE DK EE ES FI FR
    GB GR HU IE IT LI LU MC NL PL PT RO SE SI SK TR
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Original IPC: A61K-31/715(B,I,H,EP,19850101,20050415,A,L)
    A61K-39/095(B, I, H, EP, 19850101, 20050415, A, L)
    C08B-37/00(B, I, H, EP, 19740701, 20050415, A, F)
Current IPC: A61K-31/715(B,I,H,EP,19850101,20050415,A,L)
    A61K-39/095(B, I, H, EP, 19850101, 20050415, A, L)
    C08B-37/00(B, I, H, EP, 19740701, 20050415, A, F)
Current ECLA class: A61K-31/715 A61K-39/095 C08B-37/00P
Current ECLA ICO class: K61K-39:60P10
Original Abstract: Capsular saccharides derived from serogroups W135 and Y
    of ~Neisseria meningitidis~ have altered levels of O-acetylation at the
    7 and 9 positions of their sialic acid residues, and can be used to
   make immunogenic compositions. Relative to unmodifed native
    saccharides, derivatives of the invention are preferentially selected
    during conjugation to carrier proteins, and conjugates of the
    derivatives show improved immunogenicity compared to native
    polysaccharides.
Publication No. EP 2267036 Al (Update 201104 E)
Publication Date: 20101229
**Hypo- und hyperacetylierte Kapselsaccharide aus Meningokokken
  Hypo- and Hyper-Acetylated Meningococcal Capsular Saccharides
  Saccharides capsulaires de meningococcie hypo et hyperacetyles**
Assignee: Novartis Vaccines and Diagnostics S.r.l., Via Fiorentina 1, 53100
    Siena (SI), IT (NOVS)
Inventor: Costantino, Paolo, Novartis Vaccines and Diagnostics S.r.l., Via
    Fiore, I-53100, Siena, IT
  Berti, Francesco, Novartis Vaccines and Diagnostics S.r.l., Via Fiore,
    I-53100, Siena, IT
Agent: Marshall, Cameron John, Carpmaels Ransford, One Southampton Row,
   London, WC1B 5HA, GB
Language: EN
Application: EP 2010180746 A 20041004 (Local application)
  EP 2004791697 A 20041004 (Division of application)
Priority: GB 200323103 A 20031002
Related Publication: EP 1678212 A (Division of patent)
Designated States: (Regional Original) AT BE BG CH CY CZ DE DK EE ES FI FR
    GB GR HU IE IT LI LU MC NL PL PT RO SE SI SK TR
Original IPC: A61K-31/715(B,I,H,EP,20060101,20101109,A,L)
    A61K-31/715(B, I, M, 98, 20060101, 20101109, C)
    A61K-39/095(B, I, H, EP, 20060101, 20101109, A, L)
    A61K-39/095(B, I, M, 98, 20060101, 20101109, C)
    C08B-37/00(B, I, H, EP, 20060101, 20101109, A, F)
    C08B-37/00(B, I, M, 98, 20060101, 20101109, C)
Current IPC: A61K-31/715(B, I, H, EP, 20060101, 20101109, A, L)
    A61K-31/715(B, I, M, 98, 20060101, 20101109, C)
    A61K-39/095(B, I, H, EP, 20060101, 20101109, A, L)
    A61K-39/095 (B, I, M, 98, 20060101, 20101109, C)
    C08B-37/00(B, I, H, EP, 20060101, 20101109, A, F)
    C08B-37/00(B, I, M, 98, 20060101, 20101109, C)
Original Abstract: Capsular saccharides derived from serogroups W135 and Y
    of Neisseria meningitidis have altered levels of O-acetylation at the 7
    and 9 positions of their sialic acid residues, and can be used to make
    immunogenic compositions. Relative to unmodified native saccharides,
    derivatives of the invention are preferentially selected during
    conjugation to carrier proteins, and conjugates of the derivatives show
    improved immunogenicity compared to native polysaccharides.
Claim:
  1.A modified saccharide, wherein:
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* i) the saccharide is a modified serogroup W135 meningococcal capsular saccharide wherein (a) <= 29% of the sialic acid residues in the saccharide are 0-acetylated at the 7 position; and/or (b) >= 26% of the sialic acid residues in the saccharide are O-acetylated at * ii) the saccharide is a modified serogroup Y meningococcal capsular saccharide, wherein (a) <= 9% of the sialic acid residues in the saccharide are O-acetylated at the 7 position; and/or (b) >= 29% or <= 27% of the sialic acid residues in the saccharide are O-acetylated at the 9 position.

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Publication No. JP 2007507578 W (Update 200725 E)
Publication Date: 20070329
Language: JA (42 pages)
Application: JP 2006530760 A 20041004 (Local application)
  WO 2004IB3366 A 20041004 (PCT Application)
Priority: GB 200323103 A 20031002
Related Publication: WO 2005033148 A (Based on OPI patent )
Original IPC: A61K-31/715(B,I,H,JP,20060101,20070302,A,L)
    A61K-31/715(B, I, M, 98, 20060101, 20070302, C)
    A61K-39/05(B, I, H, JP, 20060101, 20070302, A, L)
    A61K-39/05(B,I,M,98,20060101,20070302,C)
    A61K-39/08(B, I, H, JP, 20060101, 20070302, A, L)
    A61K-39/08(B, I, M, 98, 20060101, 20070302, C)
    A61K-39/09(B, I, H, JP, 20060101, 20070302, A, L)
    A61K-39/09(B, I, M, 98, 20060101, 20070302, C)
    A61K-39/095(B, I, H, JP, 20060101, 20070302. A. L)
    A61K-39/095 (B, I, M, 98, 20060101, 20070302, C)
    A61K-39/10(B, I, H, JP, 20060101, 20070302, A, L)
    A61K-39/10(B, I, M, 98, 20060101, 20070302, C)
    A61K-39/102(B, I, H, JP, 20060101, 20070302, A, L)
    A61K-39/102(B, I, M, 98, 20060101, 20070302, C)
    A61K-39/125(B, I, M, 98, 20060101, 20070302, C)
    A61K-39/13(B, I, H, JP, 20060101, 20070302, A, L)
    A61K-39/29(B, I, H, JP, 20060101, 20070302, A, L)
    A61K-39/29(B, I, M, 98, 20060101, 20070302, C)
    A61P-1/00(B, I, M, 98, 20060101, 20070302, C)
    A61P-1/16(B, I, H, JP, 20060101, 20070302, A, L)
    A61P-31/00(B, I, M, 98, 20060101, 20070302, C)
    A61P-31/04(B, I, H, JP, 20060101, 20070302, A, L)
    A61P-31/14(B, I, H, JP, 20060101, 20070302, A, L)
    A61P-31/20(B, I, H, JP, 20060101, 20070302, A, L)
    A61P-37/00(B,I,M,98,20060101,20070302,C)
    A61P-37/04(B, I, H, JP, 20060101, 20070302, A, L)
    C08B-37/00(B, I, H, JP, 20060101, 20070302, A, F)
    C08B-37/00(B, I, M, 98, 20060101, 20070302, C)
Current IPC: A61K-31/715(B,I,H,JP,20060101,20070302,A,L)
    A61K-31/715(B, I, H, JP, 20060101, 20070302, C, L)
    A61K-39/05(B, I, H, JP, 20060101, 20070302, A, L)
    A61K-39/05(B, I, H, JP, 20060101, 20070302, C, L)
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    A61K-39/08(B, I, H, JP, 20060101, 20070302, C, L)
    A61K-39/09(B, I, H, JP, 20060101, 20070302, A, L)
    A61K-39/09(B, I, H, JP, 20060101, 20070302, C, L)
    A61K-39/095(B, I, H, JP, 20060101, 20070302, A, L)
    A61K-39/095(B, I, H, JP, 20060101, 20070302, C, L)
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    A61K-39/102(B,I,H,JP,20060101,20070302,C,L)
    A61K-39/125(B, I, H, JP, 20060101, 20070302, C, L)
    A61K-39/13(B, I, H, JP, 20060101, 20070302, A, L)
    A61K-39/29 (B, I, H, JP, 20060101, 20070302, A, L)
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    A61P-1/16(B, I, H, JP, 20060101, 20070302, A, L)
    A61P-31/00(B, I, H, JP, 20060101, 20070302, C, L)
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    A61P-31/20(B, I, H, JP, 20060101, 20070302, A, L)
    A61P-37/00(B, I, H, JP, 20060101, 20070302, C, L)
    A61P-37/04(B,I,H,JP,20060101,20070302,A,L)
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    C07K-14/22(B,N,H,JP,20060101,20070302,A,L)
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    C08B-37/00(B, I, H, JP, 20060101, 20070302, C, F)
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    C12N-15/09(B, N, H, JP, 20060101, 20070302, C, L)
Current ECLA class: A61K-31/715 A61K-39/095 C08B-37/00P
Current ECLA ICO class: K61K-39:60P10
Current JP F-Terms: 4B024 4C085 4C086 4C090 4C201 4H045 4B024AA01 4C086AA01
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    4C086NA14 4C086ZA75 4C086ZB09 4C086ZB33 4C086ZB35
Mexico
Publication No. MX 2006003729 Al (Update 200677 E)
Publication Date: 20060701
Assignee: CHIRON SRL (CHIR)
Inventor: COSTANTINO P
Language: ES
Application: MX 20063729 A 20060403 (Local application)
 WO 2004IB3366 A 20041004 (PCT Application)
Priority: GB 200323103 A 20031002
Related Publication: WO 2005033148 A (Based on OPI patent )
Original IPC: A61K-31/715(A) A61K-39/095(B) C08B-37/00(B)
Current IPC: A61K-31/715(B,A,I,H,MX,20060101,20060623,A,F)
    A61K-31/715(B, I, H, MX, 20060101, 20060623, C, F)
    A61K-39/095(B, I, H, MX, 20060101, 20060623, A, L)
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Current ECLA class: A61K-31/715 A61K-39/095 C08B-37/00P
Current ECLA ICO class: K61K-39:60P10
New Zealand
Publication No. NZ 546668 A (Update 200946 E)
Publication Date: 20090626
Assignee: NOVARTIS VACCINES&DIAGNOSTICS INC (NOVS)
Inventor: COSTANTINO P
Language: EN
Application: NZ 546668 A 20041004 (Local application)
 WO 2004IB3366 A 20041004 (PCT Application)
Priority: GB 200323103 A 20031002
Related Publication: WO 2005033148 A (Based on OPI patent )
Original IPC: A61K-31/715(A) A61K-39/095(B) C08B-37/00(B)
Current IPC: A61K-31/715(R,A,I,M,EP,20060101,20051008,A)
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    C08B-37/00(R,I,M,EP,20060101,20051008,C)
Current ECLA class: A61K-31/715 A61K-39/095 C08B-37/00P
Current ECLA ICO class: K61K-39:60P10
Russia
Publication No. RU 2362784 C2 (Update 200950 E)
Publication Date: 20090727
**Hypo- and hyperacetilated meningococcal capsular saccharides**
Assignee: CHIRON SRL: IT (CHIR)
Language: RU
Application: RU 2006114695 A 20041004 (Local application)
  WO 2004IB3366 A 20041004 (PCT Application)
Priority: GB 200323103 A 20031002
Related Publication: WO 2005033148 A (Based on OPI patent )
Original IPC: A61K-31/715(B,I,H,RU,20060101,20090331,A,L)
    A61K-31/715(B, I, M, 98, 20060101, 20090331, C)
    A61K-39/095(B, I, H, RU, 20060101, 20090331, A, L)
    A61K-39/095(B, I, M, 98, 20060101, 20090331, C)
    C08B-37/00(B,I,H,RU,20060101,20090331,A,F)
    C08B-37/00(B,I,M,98,20060101,20090331,C)
Current IPC: A61K-31/715(B,I,H,RU,20060101,20090909,A)
    A61K-31/715(B, I, H, RU, 20090101, 20090909, C)
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    A61K-39/095(B, I, H, RU, 20090101, 20090909, C)
    C08B-37/00(B, I, H, RU, 20060101, 20090909, A)
    C08B-37/00(B, I, H, RU, 20090101, 20090909, C)
Current ECLA class: A61K-31/715 A61K-39/095 C08B-37/00P
Current ECLA ICO class: K61K-39:60P10
Original Abstract: FIELD: chemistry. SUBSTANCE: capsular saccharides,
    obtained from serogroups W135 and Y Neisseria meningitidis, have
    changed levels of O-acetylation in positions 7 and 9 of their sialic
    acid residues. Modified meningococcal capsular saccharide of serogroup
    W135, where (a) <=29% of sialic acid residues in saccharide are
    O-acetylated in position 7; and/or (b) >=26% of sialic acid residues in
    saccharide are O-acetylated in position 9 and modified meningococcal
    capsular saccharide of serogroup Y, where a) (much less than)9% of
   sialic acid residues in saccharide are O-acetylated in position 7;
    and/or (b) >=29% or <=27% of sialic acid residues in saccharide are
    O-acetylated in position 9, can be used for creation of immunogenic
   compositions - compositions for inducing formation of antibodies in
   mammals and products of conjugation with carrier-proteins. Conjugates
   of derivatives demonstrate higher immunogenity in comparison to natural
    polysaccharides.EFFECT: obtaining capsular saccharides which
    demonstrate higher immunogenity in comparison to natural
    polysaccharides.29 cl, 6 dwg
United States
Publication No. US 20100092509 A1 (Update 201027 E)
Publication Date: 20100415
**Hypo- and Hyper- Acetylated Meningococcal Capsular Saccharides**
Assignee: Costantino, Paolo, Siena, IT Residence: IT (COST-I)
  Berti, Francesco, Siena, IT Residence: IT (BERT-I)
Inventor: Berti, Francesco, Siena, IT Residence: IT
 Costantino, Paolo, Siena, IT Residence: IT
Agent: NOVARTIS VACCINES AND DIAGNOSTICS INC., INTELLECTUAL PROPERTY-
    X100B, P.O. BOX 8097, Emeryville, CA, US
Language: EN
Application: US 2007574437 A 20070425 (Local application)
 WO 2004IB3366 A 20041004 (PCT Application)
Priority: GB 200323103 A 20031002
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Original IPC: A61K-39/095(B,I,H,US,20060101,20100415,A,L)
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   A61K-39/385(B, I, H, US, 20060101, 20100415, A, F)
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    A61P-31/04(B, I, H, US, 20060101, 20100415, A, L)
    C07H-1/00(B, I, H, US, 20060101, 20100415, A, L)
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    C07K-14/22(B, I, H, US, 20060101, 20100415, A, L)
Current IPC: A61K-31/715(R,I,M,EP,20060101,20051008,A)
    A61K-31/715(R,I,M,EP,20060101,20051008,C)
    A61K-39/095(B, I, H, US, 20060101, 20100415, A, L)
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    A61K-39/116 (B, I, H, US, 20060101, 20100415, A, L)
    A61K-39/116(B, I, H, US, 20100101, 20100415, C, L)
    A61K-39/385(B, I, H, US, 20060101, 20100415, A, F)
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    C07K-14/195(B, I, H, US, 20100101, 20100415, C, L)
    C07K-14/22(B, I, H, US, 20060101, 20100415, A, L)
    C08B-37/00(R, I, M, EP, 20060101, 20051008, A)
    C08B-37/00(R, I, M, EP, 20060101, 20051008, C)
Current ECLA class: A61K-31/715 A61K-39/095 C08B-37/00P
Current ECLA ICO class: K61K-39:60P10
Current US Class (main): 424-197110
Current US Class (secondary): 424-203100 424-250100 530-395000 536-123100
Original US Class (main): 424197.11
Original US Class (secondary): 536123.1 530395 424250.1 424203.1
Original Abstract: Capsular saccharides derived from serogroups W135 and Y
    of ~Neisseria meningitidis~ have altered levels of O-acetylation at the
    7 and 9 positions of their sialic acid residues, and can be used to
    make immunogenic compositions. Relative to unmodifed native
    saccharides, derivatives of the invention are preferentially selected
    during conjugation to carrier proteins, and conjugates of the
    derivatives show improved immunogenicity compared to native
    polysaccharides.
Claim:
**1**. A modified serogroup W135 meningococcal capsular saccharide,
        wherein: (a) >=29% of the sialic acid residues in the saccharide
        are O-acetylated at the 7 position; and/or (b) <=26% of the sialic
        acid residues in the saccharide are 0-acetylated at the 9 position.
Publication No. WO 2005033148 Al (Update 200533 B)
Publication Date: 20050414
**HYPO- AND HYPER-ACETYLATED MENINGOCOCCAL CAPSULAR SACCHARIDES
  SACCHARIDES CAPSULAIRES MENINGOCOCCIOUES HYPO ET HYPERACETYLES**
Assignee: ~(except US)~ CHIRON SRL, Vie Fiorentina 1, I-53100 Siena, IT
    Residence: IT Nationality: IT (CHIR)
  ~(only US)~ COSTANTINO, Paolo, Chiron SRL, Via Fiorentina 1, I-53100
    Siena, IT Residence: IT Nationality: IT
Inventor: COSTANTINO, Paolo, Chiron SRL, Via Fiorentina 1, I-53100 Siena,
    IT Residence: IT Nationality: IT
Agent: MARSHALL, Cameron, John, Carpmaels Ransford, 43-45 Bloomsbury
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HU IE IT KE LS LU MC MW MZ NA NL OA PL PT RO SD SE SI SK SL SZ TR TZ UG
Original IPC: C08B-37/00(A) A61K-31/715(B) A61K-39/095(B)
Current IPC: A61K-31/715(R,A,I,M,EP,20060101,20051008,A)
    A61K-31/715(R, I, M, EP, 20060101, 20051008, C)
    A61K-39/095 (R, I, M, EP, 20060101, 20051008, A)
    A61K-39/095 (R, I, M, EP, 20060101, 20051008, C)
    C08B-37/00(R, I, M, EP, 20060101, 20051008, A)
    C08B-37/00(R, I, M, EP, 20060101, 20051008, C)
Current ECLA class: A61K-31/715 A61K-39/095 C08B-37/00P
Current ECLA ICO class: K61K-39:60P10
Original Abstract: Capsular saccharides derived from serogroups W135 and Y
    of ~Neisseria meningitidis~ have altered levels of O-acetylation at the
    7 and 9 positions of their sialic acid residues, and can be used to
    make immunogenic compositions. Relative to unmodifed native
    saccharides, derivatives of the invention are preferentially selected
    during conjugation to carrier proteins, and conjugates of the
    derivatives show improved immunogenicity compared to native
    polysaccharides.
  L'invention concerne des saccharides capsulaires derives des serogroupes
    W135 et Y de ~Neisseria meningitidis~ et presentant des taux modifies
    d'acetylation en ortho au niveau des sites 7 et 9 de leurs residus
    d'acide sialique, et pouvant etre utilises pour fabriquer des
    compositions immunogenes. Par rapport aux saccharides natifs non
    modifies, les derives decrits sont de preference selectionnes
    lorsqu'ils sont conjugues a des proteines vectrices, et des conjugues
    de ces derives presentent une immunogenicite amelioree par rapport aux
    polysaccharides natifs.
 10/7/15
             (Item 15 from file: 351)
DIALOG(R)File 351:Derwent WPI
(c) 2011 Thomson Reuters. All rts. reserv.
0007899642
WPI ACC NO: 1996-251554/199625
XRAM Acc No: C1996-079594
Combined vaccine against meningitis contg. oligosaccharide conjugates - of
Haemophilus influenzae and ****Neisseria**** ****meningitidis**** serotype
Patent Assignee: CHIRON SPA (CHIR); CHIRON SRL (CHIR); NOVARTIS VACCINES
  & DIAGNOSTICS INC (NOVS); BIOCINE SPA (BIOC-N); NOVARTIS
  VACCINES&DIAGNOSTICS INC (NOVS)
Inventor: CECCARINI C; COSTANTINO P; D'ASCENZI S; DASCENZI S; GIANNOZZI A;
  NORELLI F: GIANOZZI A
Patent Family (16 patents, 20 countries)
Patent
                               Application
Number
                Kind
                     Date
                                              Kind
                               Number
                                                   Date
                                                             Update
                                              A 19951102
WO 1996014086
                A1 19960517 WO 1995IB1006
                                                             199625
EP 789587
                A1 19970820 EP 1995935550
                                              A 19951102 199738 E
                               WO 1995IB1006 A 19951102
JP 10509701 W 19980922 WO 1995IB1006 A 19951102 199848 E
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Square, London WC1A 2RA, GB Language: EN (42 pages, 6 drawings)

Priority: GB 200323103 A 20031002

TT TZ UA UG US UZ VC VN YU ZA ZM ZW

Application: WO 2004IB3366 A 20041004 (Local application)

Designated States: (National Original) AE AG AL AM AT AU AZ BA BB BG BR BW BY BZ CA CH CN CO CR CU CZ DE DK DM DZ EC EE BG ES FI GB DG EG HG MF BH HU ID IL IN IS JP KE KG KP KR KZ LC LK LR LS LT LU LV MA MD MG MK NM MW MX MZ NN NI NO NZ OM PG PH PL PT RO RU SC SD SE SG SK SL SY TJ TM TN TR

(Regional Original) AT BE BG BW CH CY CZ DE DK EA EE ES FI FR GB GH GM GR

				JP	1996515175	A	19951102		
US	6251401	В1	20010626	WO	1995IB1006	A	19951102	200138	E
				US	1997836080	A	19970501		
EP	1312377	A2	20030521	EP	1995935550	A	19951102	200334	E
				EP	200375069	A	19951102		
EP	789587	В1	20030813	EP	1995935550	A	19951102	200355	E
				WO	1995IB1006	A	19951102		
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DE	69531501	E	20030918	DE	69531501	A	19951102	200369	E
				EP	1995935550	A	19951102		
				WO	1995IB1006	A	19951102		
	2204967	Т3	20040501		1995935550	A	19951102	200431	E
JP	2007169302	A	20070705	JP	1996515175	Α	19951102	200746	E
				JΡ	200783117	Α	20070327		
JP	3989951	B2	20071010	WO	1995IB1006	Α	19951102	200768	E
				JP	1996515175	A	19951102		
EP	789587	B2	20080402	EP	1995935550	A	19951102	200825	Ε
					1995IB1006	A	19951102		
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CA	2689871	A1	19960517		2204277	A	19951102	201019	E
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EΡ	2204185	A1	20100707	EP	1995935550	Α	19951102	201045	E
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JP	2011016850	A	20110127	JP	200783117	Α	19951102	201108	E
				JP	2010236921	A	20101021		

TD 1006515175

3 100E1103

Priority Applications (no., kind, date): GB 199422096 A 19941102

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Patent Details
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Number Kind Lan Pg Dwg Filing Notes WO 1996014086 A1 EN 32 6

National Designated States, Original: CA JP US

B1 EN

A2 EN

Regional Designated States, Original: AT BE CH DE DK ES FR GB GR IE IT LU

MC NL PT SE

US 6251401

EP 1312377

EP 789587 A1 EN PCT Application W0 1995IB1006
Based on OPI patent W0 1996014086
Regional Designated States,Original: AT BE CH DE DK ES FR GB GR IE IT LI

LU MC NL PT SE JP 10509701 W JA

PCT Application W0 1995TB1006
Based on OPI patent W0 1996014086
PCT Application W0 1995TB1006
Based on OPI patent W0 1996014086
Division of application EP 1995935550

Division of patent EP 789587 Regional Designated States,Original: AT BE CH DE DK ES FR GB GR IE IT LI LU MC NL PT SE

3.0

EP 789587 B1 EN

PCT Application WO 1995IB1006 Related to application EP 200375069 Related to patent EP 1312377 Based on OPI patent WO 1996014086

Regional Designated States, Original: AT BE CH DE DK ES FR GB GR IE IT LI LU MC NL PT SE

DE 69531501 E DE

Application EP 1995935550 PCT Application WO 1995IB1006 Based on OPI patent EP 789587

ES 2204967	Т3	ES		Based on OPI patent WO 1996014086 Application EP 1995935550 Based on OPI patent EP 789587
JP 2007169302	A	JA	16	Division of application JP 1996515175
JP 3989951	В2	JA	15	PCT Application WO 1995IB1006 Previously issued patent JP 10509701
EP 789587	В2			Based on OPI patent WO 1996014086 PCT Application WO 19951B1006 Related to application EP 200375069 Related to patent EP 1312377 Based on OPI patent WO 1996014086
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CA 2204277	С	EN		PCT Application WO 1995IB1006 Based on OPI patent WO 1996014086
CA 2689871	A1	EN		Division of application CA 2204277
EP 2204185	A1	EN		Division of application EP 1995935550
				Division of application EP 200375069
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	A	JA	16	Division of application JP 200783117

Alerting Abstract WO A1

Combined meningitis vaccine comprises Hib (Haemophilus influenzae type b) and MenC (***Neisseria*** ****meningitidis**** servtype C) oligosaccharide conjugates (OC), opt. also a MenB OC. Also new are Hib.

MenC and opt. MenB OC for simultaneous, separate or sequential admin...

USE - The vaccines are used to treat or prevent bacterial meningitis. Admin is at a rate of 2-10 mug per dose, given intramuscularly at 2, 4 and 6 months of age.

ADVANTAGE - The vaccine protects against both major causes of bacterial maingitis in a single compsn. It is inexpensive and safer with no interaction between the different antigens.

Class Codes

International Classification (Main): A61K-039/02 (Additional/Secondary): A61K-039/095, A61K-039/102 International Classification (+ Attributes) IPC + Level Value Position Status Version A61K-039/02 A I F B 20060101 A61K-0039/02 A I L B 20060101 A61K-0039/095 A I F B 20060101 A61K-0039/095 A I L B 20060101

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A61K-0039/095 A I L R 20060101
 A61K-0039/102 A I
                       R 20060101
 A61K-0039/102 A I F B 20060101
 A61K-0039/102 A I L B 20060101
 A61K-0039/102 A I L R 20060101
 A61K-0039/116 A I
                       R 20060101
 A61K-0039/116 A I F B 20060101
 A61K-0039/116 A I L B 20060101
 A61K-0047/48 A I F R 20060101
 A61P-0031/04 A I L B 20060101
 A61P-0037/04 A I L B 20060101
 C07K-0014/195 A I L B 20060101
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 A61K-0039/02 C I
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 A61K-0039/02 C I F B 20100101
 A61K-0039/02 C I L B 20100101
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                          20100101
 A61K-0039/116 C
                  Т
                       R 20060101
 A61K-0039/116 C
                 I L B
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 A61P-0037/00 C I L B 20100101
 C07K-0014/195 C I L B 20060101
 C07K-0014/195 C I L R 20060101
ECLA: A61K-039/102, A61K-039/102+M, A61K-039/116
US Classification, Current Main: 424-197110; Secondary: 424-184100,
424-193100, 424-194100, 424-203100, 424-234100, 424-250100, 424-256100,
424-831000
US Classification, Issued: 424197.11, 424203.1, 424250.1, 424256.1,
 424234.1, 424184.1, 424831, 424193.1, 424194.1
JP Classification
 FI Term
                  Facet Rank Type
A61K-039/095
A61K-039/102
A61K-039/116
                         A main
A61K-047/48
A61K-047/48
                 7.
A61P-031/04
C07K-014/195
F-Term View Point Additional
Theme
       + Figure Code
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 4C085
 4C201
4H045
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4C085
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4C085

BA18

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          CC07
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          CC24
 4H045
           DA83
 4C085
           DD35
 4C085
           DD84
 4C085
           DD86
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          EA31
 4C085
          EE03
 4C085
          EE06
 4C076
          EE30
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 4C085
           GG03
 4C085
           GG 04
File Segment: CPI
DWPI Class: B04: D16
Manual Codes (CPI/A-M): B04-C02V; B04-F10A; B14-A01A5; B14-S11B; D05-H07
Original Publication Data by Authority
Canada
Publication No. CA 2204277 C (Update 201011 E)
Publication Date: 20100202
Assignee: NOVARTIS VACCINES & DIAGNOSTICS INC (NOVS)
Inventor: CECCARINI C
 COSTANTINO P
  DASCENZI S
 GIANNOZZI A
 NORELLI F
Language: EN
Application: CA 2204277 A 19951102 (Local application)
  WO 1995IB1006 A 19951102 (PCT Application)
Priority: GB 199422096 A 19941102
Related Publication: WO 1996014086 A (Based on OPI patent )
Original IPC: A61K-39/02(I,CA,20060101,A,L) A61K-39/02(I,M,98,20060101,C)
    A61K-39/095(I,CA,20060101,A,F) A61K-39/095(I,M,98,20060101,C)
    A61K-39/102(I,CA,20060101,A,L) A61K-39/102(I,M,98,20060101,C)
Current IPC: A61K-39/02(B,I,H,CA,20060101,19970804,A,L)
    A61K-39/02(B, I, H, CA, 20100101, 19970804, C, L)
    A61K-39/095(B, I, H, CA, 20060101, 19970804, A, F)
    A61K-39/095(B,I,H,CA,20100101,19970804,C,F)
    A61K-39/102(R, I, M, EP, 20060101, 20051008, A, L)
    A61K-39/102(R,I,M,EP,20100101,20051008,C,L)
    A61K-39/116 (R, I, M, EP, 20060101, 20051008, A)
    A61K-39/116(R, I, M, EP, 20060101, 20051008, C)
    A61K-47/48(R,I,M,JP,20060101,20051220,A,F)
    A61K-47/48(R,I,M,JP,20060101,20051220,C,F)
    C07K-14/195(R,I,M,JP,20060101,20051220,A,L)
   C07K-14/195(R,I,M,JP,20060101,20051220,C,L)
Current ECLA class: A61K-39/102 A61K-39/102+M A61K-39/116
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Publication No. CA 2689871 A1 (Update 201019 E)
Publication Date: 19960517
Assignee: NOVARTIS VACCINES & DIAGNOSTICS INC (NOVS)
Inventor: CECCARINI C
  COSTANTINO P
  DASCENZI S
  GIANNOZZI A
 NORELLI F
Language: EN
Application: CA 2689871 A 19951102 (Local application)
  CA 2204277 A 19951102 (Division of application)
Priority: GB 199422096 A 19941102
Original IPC: A61K-39/095(B,I,H,CA,20060101,20100215,A,F)
    A61K-39/095(B,I,H,CA,20100101,20100215,C,F)
    A61K-39/102(R,I,M,EP,20060101,20051008,A)
    A61K-39/102(R, I, M, EP, 20060101, 20051008, C)
    A61K-39/116(B, I, H, CA, 20060101, 20100215, A, L)
    A61K-39/116(B, I, H, CA, 20100101, 20100215, C, L)
    A61K-47/48(R, I, M, JP, 20060101, 20051220, A, F)
    A61K-47/48(R, I, M, JP, 20060101, 20051220, C, F)
    A61P-31/00(B,I,H,CA,20100101,20100215,C,L)
    A61P-31/04(B, I, H, CA, 20060101, 20100215, A, L)
    A61P-37/00(B, I, H, CA, 20100101, 20100215, C, L)
    A61P-37/04(B, I, H, CA, 20060101, 20100215, A, L)
    C07K-14/195(R, I, M, JP, 20060101, 20051220, A, L)
    C07K-14/195(R,I,M,JP,20060101,20051220,C,L)
Current IPC: A61K-39/095(B,I,H,CA,20060101,20100215,A,F)
    A61K-39/095(B, I, H, CA, 20100101, 20100215, C, F)
    A61K-39/102(R, I, M, EP, 20060101, 20051008, A)
    A61K-39/102(R, I, M, EP, 20060101, 20051008, C)
    A61K-39/116(B, I, H, CA, 20060101, 20100215, A, L)
    A61K-39/116(B, I, H, CA, 20100101, 20100215, C, L)
    A61K-47/48(R,I,M,JP,20060101,20051220,A,F)
    A61K-47/48(R, I, M, JP, 20060101, 20051220, C, F)
    A61P-31/00(B, I, H, CA, 20100101, 20100215, C, L)
    A61P-31/04(B, I, H, CA, 20060101, 20100215, A, L)
    A61P-37/00(B, I, H, CA, 20100101, 20100215, C, L)
    A61P-37/04(B, I, H, CA, 20060101, 20100215, A, L)
    C07K-14/195(R, I, M, JP, 20060101, 20051220, A, L)
    C07K-14/195(R,I,M,JP,20060101,20051220,C,L)
Current ECLA class: A61K-39/102 A61K-39/102+M A61K-39/116
Germany
Publication No. DE 69531501 E (Update 200369 E)
Publication Date: 20030918
Assignee: CHIRON SPA; IT (CHIR)
Language: DE
Application: DE 69531501 A 19951102 (Local application)
  EP 1995935550 A 19951102 (Application)
  WO 1995IB1006 A 19951102 (PCT Application)
Priority: GB 199422096 A 19941102
Related Publication: EP 789587 A (Based on OPI patent )
  WO 1996014086 A (Based on OPI patent )
Original IPC: A61K-39/02(A) A61K-39/02(A) A61K-39/095(B) A61K-39/095(B)
    A61K-39/102(B) A61K-39/102(B)
Current IPC: A61K-39/02(A) A61K-39/02(A) A61K-39/095(B) A61K-39/095(B)
    A61K-39/102(B) A61K-39/102(B)
Publication No. EP 1312377 A2 (Update 200334 E)
Publication Date: 20030521
```

**Kombiniertes Meningitis Vakzin

```
Combined meningitis vaccine
  Vaccin polyvalent anti-meningite**
Assignee: Chiron S.p.A., Via Fiorentina, 1, 53100 Siena, IT (CHIR)
Inventor: Ceccarini, Costante, Chrion SpA, Via Fiorentina 1, 53100 Siena,
  d'Ascenzi, Sandro, Chrion SpA, Via Fiorentina 1, 53100 Siena, IT
  Costantino, Paolo, Chrion SpA, Via Fiorentina 1, 53100 Siena, IT
 Norelli, Francesco, Chrion SpA, Via Fiorentina 1, 53100 Siena, IT
 Giannozzi, Aldo, Chrion SpA, Via Fiorentina 1, 53100 Siena, IT
Agent: Marshall, Cameron John, Carpmaels Ransford, 43 Bloomsbury Square,
    London WC1A 2RA, GB
Language: EN
Application: EP 1995935550 A 19951102 (Division of application)
  EP 200375069 A 19951102 (Local application)
Priority: GB 199422096 A 19941102
Related Publication: EP 789587 A (Division of patent)
Designated States: (Regional Original) AT BE CH DE DK ES FR GB GR IE IT LI
    LU MC NL PT SE
Original IPC: A61K-39/02(A) A61K-39/095(B) A61K-39/102(B)
Current IPC: A61K-39/095(R,I,M,JP,20060101,20051220,A,L)
    A61K-39/095(R, I, M, JP, 20060101, 20051220, C, L)
    A61K-39/102(R, I, M, EP, 20060101, 20051008, A)
    A61K-39/102(R, I, M, EP, 20060101, 20051008, C)
    A61K-39/116(R, I, M, EP, 20060101, 20051008, A)
    A61K-39/116(R, I, M, EP, 20060101, 20051008, C)
    A61K-47/48(R, I, M, JP, 20060101, 20051220, A, F)
    A61K-47/48(R, I, M, JP, 20060101, 20051220, C, F)
    C07K-14/195(R, I, M, JP, 20060101, 20051220, A, L)
    C07K-14/195(R,I,M,JP,20060101,20051220,C,L)
Current ECLA class: A61K-39/102 A61K-39/102+M A61K-39/116
Original Abstract: A combined vaccine for bacterial meningitis comprises
    Hib and MenC saccharides. div
Claim:
  1.A combination meningitis vaccine comprising Hib and MenC saccharides.
  2.A vaccine according to claim 1 wherein the Hib and/or MenC saccharide
        is an oligosaccharide.
Publication No. EP 2204185 A1 (Update 201045 E)
Publication Date: 20100707
**Kombiniertes Meningitis Vakzin
  Combined meningitis vaccine
 Vaccin polyvalent anti-meningite**
Assignee: Novartis Vaccines and Diagnostics S.r.l., Via Fiorentina 1, 53100
    Siena (SI), IT (NOVS)
Inventor: Ceccarini, Costante, Via Fiorentina 1, 53100 Siena, IT
  Costantino, Paolo, Via Fiorentina 1, 53100 Siena, IT
  DASCENZI S. IT
  Gianozzi, Aldo, Via Fiorentina 1, 53100 Siena, IT
  Norelli, Francesco, Via Fiorentina 1, 53100 Siena, IT
Agent: Marshall, Cameron John, Carpmaels Ransford, 43-45 Bloomsbury
   Square, London WC1A 2RA, GB
Language: EN
Application: EP 200375069 A 20030110 (Division of application)
  EP 1995935550 A 19951102 (Division of application)
  EP 201075114 A 19951102 (Local application)
Priority: GB 199422096 A 19941102
Related Publication: EP 1312377 A (Division of patent)
  EP 789587 A (Division of patent)
Designated States: (Regional Original) AT BE CH DE DK ES FR GB GR IE IT LI
    LU MC NL PT SE
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Original IPC: A61K-39/02(B,I,H,EP,20060101,20100520,A,F)
    A61K-39/02(B, I, M, 98, 20060101, 20100520, C)
    A61K-39/095(B, I, H, EP, 20060101, 20100520, A, L)
    A61K-39/095(B,I,M,98,20060101,20100520,C)
    A61K-39/102(B, I, H, EP, 20060101, 20100520, A, L)
    A61K-39/102(B, I, M, 98, 20060101, 20100520, C)
Current IPC: A61K-39/02(B,I,H,EP,20060101,20100520,A,F)
    A61K-39/02(B, I, H, EP, 20100101, 20100520, C, F)
    A61K-39/095(B, I, H, EP, 20060101, 20100520, A, L)
    A61K-39/095(B,I,H,EP,20100101,20100520,C,L) A61K-39/102(B,I,H,
    EP, 20060101, 20100520, A, L) A61K-39/102(B, I, H, EP, 20100101, 20100520, C, L)
    A61K-39/116(R, I, M, EP, 20060101, 20051008, A)
    A61K-39/116(R, I, M, EP, 20060101, 20051008, C)
    A61K-47/48(R, I, M, JP, 20060101, 20051220, A, F)
    A61K-47/48(R,I,M,JP,20060101,20051220,C,F)
    C07K-14/195(R, I, M, JP, 20060101, 20051220, A, L)
    C07K-14/195(R,I,M,JP,20060101,20051220,C,L)
Current ECLA class: A61K-39/102 A61K-39/102+M A61K-39/116
Original Abstract: A combined vaccine for bacterial meningitis comprises
    Hib and MenC saccharides.
Claim:
  1.A combination meningitis vaccine comprising Hib saccharide conjugate
        and MenC saccharide conjugate, wherein the MenC and/or Hib
        conjugates is/are in lyophilised form.
Publication No. EP 2204185 A8 (Update 201071 E)
Publication Date: 20101027
**Combined meningitis vaccine**
Assignee: NOVARTIS VACCINESDIAGNOSTICS INC; IT (NOVS)
Inventor: CECCARINI C, IT
  DASCENZI S, IT
  COSTANTINO P, IT
  NORELLI F, IT
  GIANOZZI A, IT
Language: EN
Application: EP 201075114 A 19951102 (Local application)
  EP 200375069 A 20030110 (Division of application)
  EP 1995935550 A 19951102 (Division of application)
Priority: GB 199422096 A 19941102
Related Publication: EP 1312377 A (Division of patent)
  EP 789587 A (Division of patent)
Designated States: (Regional Original) AT BE CH DE DK ES FR GB GR IE IT LI
    LU MC NL PT SE
Original IPC: A61K-39/02(B,I,H,EP,20060101,20100520,A,F)
    A61K-39/02(B,I,M,98,20060101,20100520,C)
    A61K-39/095(B, I, H, EP, 20060101, 20100520, A, L)
    A61K-39/095(B, I, M, 98, 20060101, 20100520, C)
    A61K-39/102(B, I, H, EP, 20060101, 20100520, A, L) A61K-39/102(B, I,
    M, 98, 20060101, 20100520, C)
Current IPC: A61K-39/02(B,I,H,EP,20060101,20100520,A,F)
    A61K-39/02(B, I, M, 98, 20060101, 20100520, C)
    A61K-39/095(B, I, H, EP, 20060101, 20100520, A, L)
    A61K-39/095(B,I,M,98,20060101,20100520,C)
    A61K-39/102(B, I, H, EP, 20060101, 20100520, A, L)
    A61K-39/102(B, I, M, 98, 20060101, 20100520, C)
Original Abstract: A combined vaccine for bacterial meningitis comprises
    Hib and MenC saccharides.
Publication No. EP 789587 A1 (Update 199738 E)
Publication Date: 19970820
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**KOMBINIERTES MENNINGITIS VAKZINE

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COMBINED MENINGITIS VACCINE
  VACCIN POLYVALENT ANTI-MENINGITE**
Assignee: BIOCINE S.p.A., Via Fiorentina, 1, I-53100 Siena, IT (BIOC-N)
  CHIRON SPA (CHIR)
Inventor: CECCARINI, Costante, Via di Cationano, 10, I-53010 Castelnuovo
    Berardenga, IT
  COSTANTINO, Paolo, Via Toscana, 11, I-53034 Colle Val d'Elsa, IT
  DASCENZI S
  NORELLI, Francesco, Via Vignale, 16, I-53100 Siena, IT
  GIANNOZZI, Aldo, Via Celso Cittadini, 3, I-53100 Siena, IT
  D'ASCENZI, Sandro, Via Palestro, 24, I-53034 Colle Val d'Elsa, IT
  D'ASCENZI, Sandro, Via Palestro, 24, I-53034 Colle Val d'Elsa, IT
  D'ASCENZI, Sandro, Via Palestro, 24, I-53034 Colle Val d'Elsa, IT
  D'ASCENZI, Sandro, Via Palestro, 24, I-53034 Colle Val d'Elsa, IT
  D'ASCENZI, Sandro, Via Palestro, 24, I-53034 Colle Val d'Elsa, IT
Agent: Hallybone, Huw George, CARPMAELS AND RANSFORD 43 Bloomsbury Square,
    London WC1A 2RA, GB
Language: EN
Application: EP 1995935550 A 19951102 (Local application)
  WO 1995IB1006 A 19951102 (PCT Application)
Priority: GB 199422096 A 19941102
Related Publication: WO 1996014086 A (Based on OPI patent )
Designated States: (Regional Original) AT BE CH DE DK ES FR GB GR IE IT LI
    LU MC NL PT SE
Original IPC: A61K-39/02(A) A61K-39/095(B) A61K-39/102(B)
Current IPC: A61K-39/095(R,A,I,M,JP,20060101,20051220,A,L)
   A61K-39/095(R,I,M,JP,20060101,20051220,C,L)
    A61K-39/102(R,I,M,EP,20060101,20051008,A)
    A61K-39/102(R, I, M, EP, 20060101, 20051008, C)
    A61K-39/116(R, I, M, EP, 20060101, 20051008, A)
    A61K-39/116(R, I, M, EP, 20060101, 20051008, C)
    A61K-47/48(R, I, M, JP, 20060101, 20051220, A, F)
    A61K-47/48(R, I, M, JP, 20060101, 20051220, C, F)
    C07K-14/195(R, I, M, JP, 20060101, 20051220, A, L)
    C07K-14/195(R, I, M, JP, 20060101, 20051220, C, L)
Current ECLA class: A61K-39/102 A61K-39/102+M A61K-39/116
Original Abstract: A combined vaccine for bacterial meningitis comprises
    Hib and MenC oligosaccharide conjugates.
Claim: Combined meningitis vaccine comprises Hib (Haemophilus influenzae
    type b) and MenC (Neisseria meningitidis serotype C) oligosaccharide
    conjugates (OC), opt. also a MenB OC. Also new are Hib. MenC and opt.
    MenB OC for simultaneous, separate or sequential admin..
Publication No. EP 789587 B1 (Update 200355 E)
Publication Date: 20030813
**KOMBINIERTES MENNINGITIS VAKZINE
  COMBINED MENINGITIS VACCINE
  VACCIN POLYVALENT ANTI-MENINGITE**
Assignee: Chiron S.p.A., Via Fiorentina, 1, 53100 Siena, IT (CHIR)
Inventor: CECCARINI, Costante, Via di Catignano, 10, I-53010 Castelnuovo
    Berardenga, IT
  COSTANTINO, Paolo, Via Toscana, 11, I-53034 Colle Val d'Elsa, IT
  D'ASCENZI, Sandro, Via Palestro, 24, I-53034 Colle Val d'Elsa, IT
 NORELLI, Francesco, Via Vignale, 16, I-53100 Siena, IT
  GIANNOZZI, Aldo, Via Celso Cittadini, 3, I-53100 Siena, IT
Agent: Hallybone, Huw George, Carpmaels and Ransford, 43 Bloomsbury Square,
   London WC1A 2RA, GB
Language: EN
Application: EP 1995935550 A 19951102 (Local application)
  WO 1995IB1006 A 19951102 (PCT Application)
  EP 200375069 A 19951102 (Related to application)
Priority: GB 199422096 A 19941102
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Related Publication: EP 1312377 A (Related to patent)
  WO 1996014086 A (Based on OPI patent )
Designated States: (Regional Original) AT BE CH DE DK ES FR GB GR IE IT LI
    LU MC NL PT SE
Original IPC: A61K-39/02(A) A61K-39/095(B) A61K-39/102(B)
Current IPC: A61K-39/095(R,I,M,JP,20060101,20051220,A,L)
    A61K-39/095(R, I, M, JP, 20060101, 20051220, C, L)
    A61K-39/102(R, I, M, EP, 20060101, 20051008, A)
    A61K-39/102(R,I,M,EP,20060101,20051008,C)
    A61K-39/116(R, I, M, EP, 20060101, 20051008, A)
   A61K-39/116(R,I,M,EP,20060101,20051008,C)
    A61K-47/48(R, I, M, JP, 20060101, 20051220, A, F)
   A61K-47/48(R, I, M, JP, 20060101, 20051220, C, F)
    C07K-14/195(R,I,M,JP,20060101,20051220,A,L)
    C07K-14/195(R,I,M,JP,20060101,20051220,C,L)
Current ECLA class: A61K-39/102 A61K-39/102+M A61K-39/116
Claim.
  1. Meningitis-Kombinationsimpfstoff, umfassend Hib- und
        MenC-Oligosaccharid-Konjugate.
  2. Impfstoff nach Anspruch 1, weiterhin umfassend ein
        MenB-Oligosaccharid-Konjugat.
  1.A combination meningitis vaccine comprising Hib and MenC
        oligosaccharide conjugates.
  2.A vaccine according to claim 1 further comprising a MenB
        oligosaccharide conjugate.
  1. Vaccin polyvalent anti-meningite comprenant des conjugues
        d'oligosaccharide Hib et MenC.
Publication No. EP 789587 B2 (Update 200825 E)
Publication Date: 20080402
**KOMBINIERTES MENNINGITIS VAKZINE
  COMBINED MENINGITIS VACCINE
  VACCIN POLYVALENT ANTI-MENINGITE**
Assignee: Novartis Vaccines and Diagnostics S.r.l., Via Fiorentina 1, 53100
    Siena (SI), IT (NOVS)
Inventor: CECCARINI, Costante, Via di Catignano, 10, I-53010 Castelnuovo
    Berardenga, IT
  COSTANTINO, Paolo, Via Toscana, 11, I-53034 Colle Val d'Elsa, IT
  DASCENZI S
 NORELLI, Francesco, Via Vignale, 16, I-53100 Siena, IT
  GIANNOZZI, Aldo, Via Celso Cittadini, 3, I-53100 Siena, IT
Agent: Hallybone, Huw George, Carpmaels and Ransford, 43-45 Bloomsbury
    Square, London WC1A 2RA, GB
Language: EN
Application: EP 1995935550 A 19951102 (Local application)
  WO 1995IB1006 A 19951102 (PCT Application)
  EP 200375069 A 20030110 (Related to application)
Priority: GB 199422096 A 19941102
Related Publication: EP 1312377 A (Related to patent)
  WO 1996014086 A (Based on OPI patent )
Designated States: (Regional Original) AT BE CH DE DK ES FR GB GR IE IT LI
    LU MC NL PT SE
Original IPC: A61K-39/02(B,I,H,EP,20060101,19960625,A,F)
    A61K-39/02(B, I, M, 98, 20060101, 19960625, C)
    A61K-39/095(B, I, H, EP, 20060101, 19960625, A, L)
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A61K-39/095(B, I, M, 98, 20060101, 19960625, C)
    A61K-39/102(B, I, H, EP, 20060101, 19960625, A, L)
    A61K-39/102(B, I, M, 98, 20060101, 19960625, C)
Current IPC: A61K-39/02(B,I,H,EP,20060101,19960625,A,F)
    A61K-39/02(B, I, H, EP, 20060101, 19960625, C, F)
    A61K-39/095(B, I, H, EP, 20060101, 19960625, A, L)
    A61K-39/095(B, I, H, EP, 20060101, 19960625, C, L)
    A61K-39/102(B, I, H, EP, 20060101, 19960625, A, L)
    A61K-39/102(B, I, H, EP, 20060101, 19960625, C, L)
    A61K-39/116(R, I, M, EP, 20060101, 20051008, A)
    A61K-39/116(R,I,M,EP,20060101,20051008,C)
    A61K-47/48(R, I, M, JP, 20060101, 20051220, A, F)
    A61K-47/48(R, I, M, JP, 20060101, 20051220, C, F)
    C07K-14/195(R,I,M,JP,20060101,20051220,A,L)
    C07K-14/195(R,I,M,JP,20060101,20051220,C,L)
Current ECLA class: A61K-39/102 A61K-39/102+M A61K-39/116
Claim.
  1.Meningitis-Kombinationsimpfstoff, umfassend Hib- und
        MenC-Oligosaccharid-Konjugate.
  1.A combination meningitis vaccine comprising Hib and MenC
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oligosaccharide conjugates.

 Vaccin polyvalent anti-meningite comprenant des conjugues d'oligosaccharide Hib et MenC.

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Spain
Publication No. ES 2204967 T3 (Update 200431 E)
Publication Date: 20040501
Assignee: CHIRON SRL (CHIR)
Language: ES
Application: EP 1995935550 A 19951102 (Application)
Priority: GB 199422096 A 19941102
Related Publication: EP 789587 A (Based on OPI patent )
Original IPC: A61K-39/02(A) A61K-39/095(B) A61K-39/102(B)
Current IPC: A61K-39/095(R,I,M,JP,20060101,20051220,A,L)
    A61K-39/095(R,I,M,JP,20060101,20051220,C,L)
    A61K-39/102(R,I,M,EP,20060101,20051008,A)
    A61K-39/102(R, I, M, EP, 20060101, 20051008, C)
   A61K-39/116(R, I, M, EP, 20060101, 20051008, A)
    A61K-39/116(R, I, M, EP, 20060101, 20051008, C)
   A61K-47/48(R, I, M, JP, 20060101, 20051220, A, F)
   A61K-47/48(R,I,M,JP,20060101,20051220,C,F)
    C07K-14/195(R,I,M,JP,20060101,20051220,A,L)
    C07K-14/195(R, I, M, JP, 20060101, 20051220, C, L)
Current ECLA class: A61K-39/102 A61K-39/102+M A61K-39/116
Publication No. JP 10509701 W (Update 199848 E)
Publication Date: 19980922
Assignee: BIOCINE SPA (BIOC-N)
Inventor: CECCARINI C
 COSTANTINO P
  DASCENZI S
 NORELLI F
 GIANNOZZI A
Language: JA (30 pages)
Application: WO 1995IB1006 A 19951102 (PCT Application)
  JP 1996515175 A 19951102 (Local application)
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Priority: GB 199422096 A 19941102
Related Publication: WO 1996014086 A (Based on OPI patent )
Original IPC: A61K-39/102(A) A61K-39/095(B) A61K-47/48(B) C07K-14/195(B)
Current IPC: A61K-39/095(R,A,I,M,JP,20060101,20051220,A,L)
    A61K-39/095(R, I, M, JP, 20060101, 20051220, C, L)
    A61K-39/102(R, I, M, EP, 20060101, 20051008, A)
    A61K-39/102(R, I, M, EP, 20060101, 20051008, C)
    A61K-39/116(R,I,M,EP,20060101,20051008,A)
    A61K-39/116(R, I, M, EP, 20060101, 20051008, C)
    A61K-47/48(R,I,M,JP,20060101,20051220,A,F)
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    C07K-14/195(R,I,M,JP,20060101,20051220,A,L)
    C07K-14/195(R, I, M, JP, 20060101, 20051220, C, L)
Current ECLA class: A61K-39/102 A61K-39/102+M A61K-39/116
Publication No. JP 2007169302 A (Update 200746 E)
Publication Date: 20070705
**COMBINED MENINGITIS VACCINE**
Assignee: CHIRON SRL (CHIR-N)
Inventor: CECCARINI COSTANTE
  COSTANTINO PAOLO
  D'ASCENZI SANDRO
 NORELLI FRANCESCO
 GIANNOZZI ALDO
Language: JA (16 pages)
Application: JP 1996515175 A 19951102 (Division of application)
  JP 200783117 A 20070327 (Local application)
Priority: GB 199422096 A 19941102
Original IPC: A61K-39/095(B,I,H,JP,20060101,20070608,A,L)
    A61K-39/095(B, I, M, 98, 20060101, 20070608, C)
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    A61P-31/00(B, I, M, 98, 20060101, 20070608, C)
    A61P-31/04(B, I, H, JP, 20060101, 20070608, A, L)
Current IPC: A61K-39/095(B,I,H,JP,20060101,20070608,A,L)
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    A61K-39/102(B, I, H, JP, 20060101, 20070608, A, F)
    A61K-39/102(B, I, H, JP, 20060101, 20070608, C, F)
    A61K-39/116 (R, I, M, EP, 20060101, 20051008, A)
    A61K-39/116(R, I, M, EP, 20060101, 20051008, C)
    A61K-47/48(R, I, M, JP, 20060101, 20051220, A, F)
    A61K-47/48(R, I, M, JP, 20060101, 20051220, C, F)
    A61P-31/00(B, I, H, JP, 20060101, 20070608, C, L)
    A61P-31/04(B, I, H, JP, 20060101, 20070608, A, L)
    C07K-14/195(R, I, M, JP, 20060101, 20051220, A, L)
    C07K-14/195(R, I, M, JP, 20060101, 20051220, C, L)
Current ECLA class: A61K-39/102 A61K-39/102+M A61K-39/116
Current JP F-Terms: 4C085 4C201 4C085BA16 4C085BA18 4C085CC07 4C085EE03
    4C085GG01 4C085GG03
Claim: A combination meningitis vaccine which is described in
    this-application specification.
Publication No. JP 2011016850 A (Update 201108 E)
Publication Date: 20110127
**Combination meningitis vaccine**
Assignee: CHIRON SPA; JP (CHIR)
Language: JA (16 pages)
Application: JP 2010236921 A 20101021 (Local application)
  JP 200783117 A 19951102 (Division of application)
Priority: GB 199422096 A 19941102
Original IPC: A61K-39/116(B,I,H,JP,20060101,20101224,A,F)
Current IPC: A61K-39/116(B,I,H,JP,20060101,20101224,A,F)
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Current ECLA class: A61K-39/102 A61K-39/102+M A61K-39/116
Current JP FI-Terms: A61K-39/116 (main, A)
Current JP F-Terms: 4C085 4C085AA04 4C085BA16 4C085BA18 4C085BB24 4C085CC07
    4C085EE06 4C085FF14
Original Abstract: It is providing Hib and a meningococcus combination
    vaccine. This is used in prevention of bacterial meningitis and should
    make an economical, safe, and convenient vaccination possible with
    respect to the leading cause of a meningitis. In the combination vaccine
    for treatment of bacterial meningitis, especially one embodiment. The
    combination vaccine etc. which are effectively protected from the
    infection by the Haemophilus influenzae B type | mold (Hib), the
   Neisseria meningitidis (meningococcus) B serotype, and C serotype
    (MenB, MenC). AbsenceThis invention relates to the combination vaccine
    for treatment of bacterial meningitis. Especially a combination vaccine
    is effectively protected from the infection by the Haemophilus
    influenzae B type | mold (Hib), the Neisseria meningitidis
    (meningococcus) B serotype, and C serotype (MenB, MenC).
Claim: Invention as described in a specification.
Publication No. JP 3989951 B2 (Update 200768 E)
Publication Date: 20071010
**Combination meningitis vaccine**
Assignee: CHIRON SPA; JP (CHIR-N)
Language: JA (15 pages)
Application: WO 1995IB1006 A 19951102 (PCT Application)
  JP 1996515175 A 19951102 (Local application)
Priority: GB 199422096 A 19941102
Related Publication: JP 10509701 A (Previously issued patent)
 WO 1996014086 A (Based on OPI patent )
Original IPC: A61K-39/095(B,I,H,JP,20060101,20070920,A,L)
    A61K-39/095(B, I, M, 98, 20060101, 20070920, C)
    A61K-39/102(B, I, H, JP, 20060101, 20070920, A, F)
    A61K-39/102(B, I, M, 98, 20060101, 20070920, C)
    A61K-47/48(B, I, H, JP, 20060101, 20070920, A, L)
    A61K-47/48(B, I, M, 98, 20060101, 20070920, C)
    C07K-14/195(B, I, H, JP, 20060101, 20070920, A, L)
    C07K-14/195(B, I, M, 98, 20060101, 20070920, C)
Current IPC: A61K-39/095(B,I,H,JP,20060101,20070920,A,L)
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    A61K-39/102(B, I, H, JP, 20060101, 20070920, A, F)
    A61K-39/102(B, I, H, JP, 20060101, 20070920, C, F)
    A61K-39/116 (R, I, M, EP, 20060101, 20051008, A)
    A61K-39/116(R, I, M, EP, 20060101, 20051008, C)
    A61K-47/48(R, I, M, JP, 20060101, 20051220, A, F)
    A61K-47/48(R, I, M, JP, 20060101, 20051220, C, F)
    C07K-14/195(B, I, H, JP, 20060101, 20070920, A, L)
    C07K-14/195(B, I, H, JP, 20060101, 20070920, C, L)
Current ECLA class: A61K-39/102 A61K-39/102+M A61K-39/116
Current JP F-Terms: 4C076 4C085 4H045 4C085AA04 4H045AA30 4C085BA10
    4C085BA12 4C085BA16 4C085BA17 4C085BA18 4H045BA53 4C076BB15 4C085BB24
    4H045CA11 4C076CC06 4C085CC21 4C085CC24 4H045DA83 4C085DD35 4C085DD84
    4C085DD86 4H045EA31 4C085EE03 4C076EE30 4C076EE59 4C076FF68 4C085GG03
Claim: The combination meningitis vaccine containing a Hib oligosaccharide
    bonded material and a MenC olicosaccharide bonded material.
United States
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Publication No. US 6251401 B1 (Update 200138 E)

Inventor: Ceccarini, Costante, Castelnuovo Berardenga, IT

Publication Date: 20010626 **Combined meningitis vaccine.** Assignee: Chiron S.p.A., Siena, IT (CHIR)

```
Costantino, Paolo, Colle Val DprimeElsa, IT
  DprimeAscenzi, Sandro, Colle Val DprimeElsa, IT
  Norelli, Francesco, Siena, IT
  Giannozzi, Aldo, Siena, IT
Agent: Trujillo; Doreen Y.
  Harbin: Alisa A.
  Blackburn; Robert P.
Language: EN
Application: WO 1995IB1006 A 19951102 (PCT Application)
 US 1997836080 A 19970501 (Local application)
Priority: GB 199422096 A 19941102
Related Publication: WO 1996014086 A (Based on OPI patent )
Original IPC: A61K-39/385(A) A61K-39/095(B) A61K-39/102(B) A61K-39/116(B)
Current IPC: A61K-39/095(R,A,I,M,JP,20060101,20051220,A,L)
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    A61K-39/102(R, I, M, EP, 20060101, 20051008, A)
    A61K-39/102(R, I, M, EP, 20060101, 20051008, C)
    A61K-39/116(R, I, M, EP, 20060101, 20051008, A)
    A61K-39/116(R, I, M, EP, 20060101, 20051008, C)
    A61K-47/48(R, I, M, JP, 20060101, 20051220, A, F)
    A61K-47/48(R,I,M,JP,20060101,20051220,C,F)
    C07K-14/195(R,I,M,JP,20060101,20051220,A,L)
    C07K-14/195(R, I, M, JP, 20060101, 20051220, C, L)
Current ECLA class: A61K-39/102 A61K-39/102+M A61K-39/116
Current US Class (main): 424-197110
Current US Class (secondary): 424-184100 424-193100 424-194100 424-203100
    424-234100 424-250100 424-256100 424-831000
Original US Class (main): 424197.11
Original US Class (secondary): 424203.1 424250.1 424256.1 424234.1 424184.1
    424831 424193.1 424194.1
Original Abstract: A combined vaccine for bacterial meningitis comprises
    Hib and MenC oligosaccharide conjugates.
Claim:
  1.A combination bacterial meningitis vaccine comprising
~Haemophilus
        influenzae~type B and ~Neisseria meningitidis ~serotype C capsular
        oligosaccharide conjugates, wherein capsular oligosaccharides of
        ~Haemophilus influenzae~type B and ~Neisseria meningitidis
        ~serotype C are size-selected in order to exclude short-chain
        oligomers having a degree of polymerisation of less than 4.
Publication No. WO 1996014086 Al (Update 199625 B)
Publication Date: 19960517
**COMBINED MENINGITIS VACCINE**
Assignee: BIOCINE S.P.A., IT (BIOC-N)
Inventor: CECCARINI, COSTANTE, IT
  COSTANTINO, PAOLO, IT
  DASCENZI S
  NORELLI, FRANCESCO, IT
  GIANNOZZI, ALDO, IT
  D'ASCENZI, SANDRO, IT
Language: EN (32 pages, 6 drawings)
Application: WO 1995IB1006 A 19951102 (Local application)
Priority: GB 199422096 A 19941102
Designated States: (National Original) CA JP US
  (Regional Original) AT BE CH DE DK ES FR GB GR IE IT LU MC NL PT SE
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Original IPC: A61K-39/02(A) A61K-39/095(B) A61K-39/102(B)
Current IPC: A61K-39/095(R, A, I, M, JP, 20060101, 20051220, C, L)
A61K-39/095(R, I, M, JP, 20060101, 20051220, C, L)
A61K-39/102(R, I, M, EP, 20060101, 20051008, A)
A61K-39/102(R, I, M, EP, 20060101, 20051008, C)
A61K-39/101(R, I, M, EP, 20060101, 20051008, A)
A61K-39/116(R, I, M, EP, 20060101, 20051008, C)
A61K-47/48(R, I, M, JP, 20060101, 20051220, A, F)
A61K-47/48(R, I, M, JP, 20060101, 20051220, A, F)
C07K-14/195(R, I, M, JP, 20060101, 20051220, A, L)
C07K-14/195(R, I, M, JP, 20060101, 20051220, C, L)
Current ECLE Class: A61K-39/102 A61K-39/102+M
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Original Abstract: A combined vaccine for bacterial meningitis comprises Hib and MenC oligosaccharide conjugates.

10/7/16 (Item 16 from file: 351) DIALOG(R)File 351:Derwent WPI

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0006496336

WPI ACC NO: 1993-303147/199338

XRAM Acc No: C1993-134992

New conjugates of heat shock protein and oligo- or ****polysaccharide**** - used in vaccines or to prevent or treat bacterial infection

Patent Assignee: BIOCINE SCLAVO SPA (ISTS); BIOCINE SPA (ISTS); CHIRON SPA (CHIR); IST RICERCHE IMMUNOBIOLOGICHE SIENA SRL (RICE-M) Inventor: CONSTANTINO P; COSTANTINO P; NORELLI P; RAPPUOLI R; VITI S

Patent Family (15 patents, 41 countries)

Patent		Application								
Number		Kind	Date	Number		Kind	Date	Update		
		1993017712	A2	19930916		1993EP516	A	19930308	199338	В
		199337462	A	19931005		199337462	A	19930308	199405	E
		632727	A1	19950111		1993906489	A	19930308	199507	E
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	T-T-O	1993017712	A3	19931111		1993EP516	A	19930308	199514	Е
		7504423	W	19950518		1993515333	A	19930308	199528	E
	JE	1304423	99	19930310		1993EP516	A	19930308	199320	E
	TT	1262896	В	19960722		1992FI58	A	19920306	199709	Е
		632727	B1	19971229		1993906489	A	19930308	199805	E
	EP	632121	ы	199/1229		1993906489 1993EP516	A	19930308	199805	E
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	DE	69315993	E	19980205		69315993	A	19930308	199811	Ε
						1993906489	A	19930308		
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	US	6403099	B1	20020611		1993EP516	A	19930308	200244	E
						1994256847	A	19941101		
	CA	2131551	С	20030520		2131551	A	19930308	200335	Ε
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	JΡ	2004346083	A	20041209		1993515333	A	19930308	200481	Ε
						2004216652	A	20040723		
	JΡ	2005068131	A	20050317		1993515333	A	19930308	200520	E
					JP	2004137928	A	20040506		
	JΡ	3641483	B2	20050420	JP	1993515333	A	19930308	200527	E
					WO	1993EP516	A	19930308		
	JΡ	2009102344	A	20090514	JP	2004137928	A	19930308	200933	E
					JP	2008312770	A	20081208		
	JΡ	2011052000	A	20110317	JP	2008312770	A	19930308	201121	E
					JP	2010239145	A	20101025		

Priority Applications (number, kind, date): IT 1992FI58 A 19920306

Patent Details

AU 199337462 A EN Based on OPI patent WO 1993017712
EP 632727 Al EN PCT Application WO 1993B916
Based on OPI patent WO 1993017712
Regional Designated States,Original: AT BE CH DE DK ES FR GB GR IE IT LI
LU MC NL PT SE
WO 1993017712 A3 EN

 JP 7504423
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 PCT Application Wo 1993EP516

 EP 632727
 B1
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 27
 PCT Application Wo 1993EP516

 Based on OPI patent
 WO 1993EP516

 Based on OPI patent
 WO 1993UP7712

Regional Designated States, Original: AT BE CH DE DK ES FR GB GR IE IT LI LU MC NL PT SE

DE 69315993 DE Application EP 1993906489 PCT Application WO 1993EP516 Based on OPI patent EP 632727 Based on OPI patent WO 1993017712 US 6403099 B1 EN PCT Application WO 1993EP516 Based on OPI patent WO 1993017712 CA 2131551 PCT Application WO 1993EP516 ΕN Based on OPI patent WO 1993017712 JP 2004346083 Division of application JP 1993515333 ·TΑ 36 JP 2005068131 JA 35 Division of application JP 1993515333 JP 3641483 B2 JA 27 PCT Application WO 1993EP516 Previously issued patent JP 07504423 Based on OPI patent WO 1993017712 JP 2009102344 JA 35 Division of application JP 2004137928 JP 2011052000 JA 35 Division of application JP 2008312770

Alerting Abstract WO A2

A conjugate cpd. comprises at least one heat shock protein (hsp) or portion including at least one immunostimulatory domain; and at least one oligosaccharide or ****polysaccharide****. The shp may be e.g. M. bovis BCG GroEl-type 65 kD hsp (hsp R65), recombinant M. tuberculosis Dnak-type 60 kD hsp (hsp R70) or a hsp from H. pylori.

ÜSE/ADVANTAGE - The hops are highly conserved across bacteria and they stimulate the cellular immune system. When they are conjugated to a ****polysaccharide**** they provide an immunostimulatory effect and produce anti-****polysaccharide**** antibodies in the absence of adjuvants and of pre-sensitisation. The conjugates can be used as vaccines for prophylactic or therapuetic use. In partic, they can be used for vaccination against bacteis such as H. influenzae, Streptococcus, Salmonella and Meningococci

Title Terms/Index Terms/Additional Words: NEW; CONJUGATE; HEAT; SHOCK; PROTEIN; OLIGO; POLY; SACCHARIDE; VACCINE; PREVENT; TREAT; BACTERIA; INFECT

Class Codes
International Classification (Main): A61K-047/48
(Additional/Secondary): A61K-039/385, C07K-014/195, C07K-002/00
International Classification (+ Attributes)

IPC + Level Value Position Status Version

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A61K-0038/00 A N R 20060101
  A61K-0039/00 A I F B 20060101
 A61K-0039/00 A I L R 20060101
 A61K-0039/095 A I R 20060101
 A61K-0039/385 A I L R 20060101
 A61K-0047/48 A I R 20060101
  A61P-0031/04 A I L B 20060101
 A61P-0031/04 A I L R 20060101
 A61P-0037/04 A I L R 20060101
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C07K-0014/35 A I R 20060101
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  C07K-0014/41 A I L R 20060101
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  C12N-0015/09 A I F R 20060101
  C12P-0021/02 A I L R 20060101
  A61K-0031/715 A I L B 20060101
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 C12N-0015/09 C I F R 20060101
 C12P-0021/02 C I L R 20060101
ECLA: A61K-039/095, A61K-047/48R2V, C07K-014/205, C07K-014/35
ICO: K61K-038:00, K61K-039:00, M07K-207:00, M07K-319:35, M07K-319:40
US Classification, Issued: 424192.1, 530395, 514569, 4357.32, 43512,
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JP Classification
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A61K-031/715
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C12P-021/02
C12P-021/02
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F-Term View Point Additional
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CA05

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4B024

4B024

HA08

HA09

Α

Α

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          ZB35
File Segment: CPI
DWPI Class: B04; D16
Manual Codes (CPI/A-M): B02-V02; B04-B02B1; B04-B04A5; B04-C02; B04-D01;
 B12-A01; B12-A06; D05-C11; D05-H07
Original Publication Data by Authority
Australia
Publication No. AU 199337462 A (Update 199405 E)
Publication Date: 19931005
Assignee: BIOCINE SCLAVO SPA (ISTS)
Inventor: RAPPUOLI R
 COSTANTINO P
 VITI S
 NORELLI F
Language: EN
Application: AU 199337462 A 19930308 (Local application)
Priority: IT 1992FI58 A 19920306
Related Publication: WO 1993017712 A (Based on OPI patent )
Original IPC: A61K-47/48(A) C07K-15/00(B)
Current IPC: A61K-47/48(R,A,I,M,EP,20060101,20051206,A)
   A61K-47/48(R.I.M.EP.20060101.20051206.C)
Canada
Publication No. CA 2131551 C (Update 200335 E)
Publication Date: 20030520
Assignee: CHIRON SPA (CHIR-N)
Inventor: VITI S
 NORELLI F
  RAPPUOLI R
  COSTANTINO P
Language: EN
Application: CA 2131551 A 19930308 (Local application)
 WO 1993EP516 A 19930308 (PCT Application)
Priority: IT 1992FI58 A 19920306
Related Publication: WO 1993017712 A (Based on OPI patent )
Original IPC: A61K-39/39(A) A61K-39/02(B)
Current IPC: A61K-38/00(R,A,N,M,EP,20060101,20051008,A)
    A61K-38/00(R,N,M,EP,20060101,20051008,C)
    A61K-39/00(R,I,M,JP,20060101,20051220,A,L)
    A61K-39/00(R,I,M,JP,20060101,20051220,C,L)
    A61K-39/095(R,I,M,EP,20060101,20051008,A)
    A61K-39/095(R, I, M, EP, 20060101, 20051008, C)
    A61K-39/385(R,I,M,JP,20060101,20051220,A,L)
    A61K-39/385(R,I,M,JP,20060101,20051220,C,L)
    A61K-47/48(R, I, M, EP, 20060101, 20051008, A)
    A61K-47/48(R,I,M,EP,20060101,20051008,C)
    A61P-31/00(R, I, M, JP, 20060101, 20051220, C, L)
    A61P-31/04(R,I,M,JP,20060101,20051220,A,L)
    A61P-37/00(R,I,M,JP,20060101,20051220,C,L)
    A61P-37/04(R,I,M,JP,20060101,20051220,A,L)
    C07K-14/195(R,I,M,JP,20060101,20051220,A,L)
    C07K-14/195(R, I, M, EP, 20060101, 20051008, C)
   C07K-14/205(R, I, M, EP, 20060101, 20051008, A)
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C07K-14/35(R, I, M, EP, 20060101, 20051008, A)
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    C07K-14/41(R,I,M,JP,20060101,20051220,C,L)
    C07K-19/00(R,I,M,JP,20060101,20051220,A,L)
    C07K-19/00(R, I, M, JP, 20060101, 20051220, C, L)
    C08B-37/00(R, I, M, JP, 20060101, 20051220, A, L)
    C08B-37/00(R,I,M,JP,20060101,20051220,C,L)
    C12N-15/09(R, I, M, JP, 20060101, 20051220, A, F)
    C12N-15/09(R,I,M,JP,20060101,20051220,C,F)
    C12P-21/02(R,I,M,JP,20060101,20051220,A,L)
    C12P-21/02(R, I, M, JP, 20060101, 20051220, C, L)
Current ECLA class: A61K-39/095 A61K-47/48R2V C07K-14/205 C07K-14/35
Current ECLA ICO class: K61K-38:00 K61K-39:00 M07K-207:00 M07K-319:35
   M07K-319:40
Germany
Publication No. DE 69315993 E (Update 199811 E)
Publication Date: 19980205
Assignee: BIOCINE SPA; IT (BIOC-N)
Language: DE
Application: DE 69315993 A 19930308 (Local application)
  EP 1993906489 A 19930308 (Application)
  WO 1993EP516 A 19930308 (PCT Application)
Priority: IT 1992FI58 A 19920306
Related Publication: EP 632727 A (Based on OPI patent )
 WO 1993017712 A (Based on OPI patent )
Original IPC: A61K-47/48(A) C07K-2/00(B)
Current IPC: A61K-47/48(A) C07K-2/00(B)
Publication No. EP 632727 A1 (Update 199507 E)
Publication Date: 19950111
**KONJUGATE AUS HITZESCHOCKPROTEINEN UND OLIGO- ODER POLYSACCHARIDEN
  CONJUGATES FORMED FROM HEAT SHOCK PROTEINS AND OLIGO- OR POLYSACCHARIDES
  COMPOSES CONJUGUES OBTENUS A PARTIR DE PROTEINES DU CHOC THERMIQUE ET
    D'OLIGOSACCHARIDES OU DE POLYSACCHARIDES**
Assignee: BIOCINE SpA, Via Fiorentina, 1, I-53100 Siena, IT (ISTS)
Inventor: RAPPUOLI, Rino, Via Calamadrei, 39, Quercegrossa, I-53010
   Monteriggioni, IT
  COSTANTINO, Paolo, Via Toscano, 11, I-53034 Colle Val d'Elsa, IT
  VITI, Stefano, Via Dante Alighieri, I-53018 Sovicille, IT
 NORELLI, Francesco, Via Vignali, 16, I-53100 Siena, IT
Agent: Hallybone, Huw George, CARPMAELS AND RANSFORD 43 Bloomsbury Square,
    London WC1A 2RA, GB
Language: EN
Application: EP 1993906489 A 19930308 (Local application)
  WO 1993EP516 A 19930308 (PCT Application)
Priority: IT 1992FI58 A 19920306
Related Publication: WO 1993017712 A (Based on OPI patent )
Designated States: (Regional Original) AT BE CH DE DK ES FR GB GR IE IT LI
    LU MC NL PT SE
Original IPC: A61K-47/48(A) C07K-15/00(B)
Current IPC: A61K-38/00(R,A,N,M,EP,20060101,20051008,A)
    A61K-38/00(R,N,M,EP,20060101,20051008,C)
    A61K-39/00(R, I, M, JP, 20060101, 20051220, A, L)
    A61K-39/00(R, I, M, JP, 20060101, 20051220, C, L)
    A61K-39/095(R,I,M,EP,20060101,20051008,A)
   A61K-39/095(R,I,M,EP,20060101,20051008,C)
   A61K-39/385(R, I, M, JP, 20060101, 20051220, A, L)
   A61K-39/385(R,I,M,JP,20060101,20051220,C,L)
   A61K-47/48(R,I,M,EP,20060101,20051008,A)
   A61K-47/48(R,I,M,EP,20060101,20051008,C)
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A61P-31/00(R, I, M, JP, 20060101, 20051220, C, L)
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    C07K-14/195(R, I, M, JP, 20060101, 20051220, A, L)
    C07K-14/195(R, I, M, EP, 20060101, 20051008, C)
   C07K-14/205(R, I, M, EP, 20060101, 20051008, A)
    C07K-14/35(R, I, M, EP, 20060101, 20051008, A)
    C07K-14/41(R,I,M,JP,20060101,20051220,A,L)
    C07K-14/41(R,I,M,JP,20060101,20051220,C,L)
    C07K-19/00(R,I,M,JP,20060101,20051220,A,L) C07K-19/00(R,I,
    M, JP, 20060101, 20051220, C, L) C08B-37/00 (R, I, M, JP, 20060101, 20051220, A, L)
    C08B-37/00(R,I,M,JP,20060101,20051220,C,L)
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    C12N-15/09(R,I,M,JP,20060101,20051220,C,F)
    C12P-21/02(R, I, M, JP, 20060101, 20051220, A, L)
    C12P-21/02(R, I, M, JP, 20060101, 20051220, C, L)
Current ECLA class: A61K-39/095 A61K-47/48R2V C07K-14/205 C07K-14/35
Current ECLA ICO class: K61K-38:00 K61K-39:00 M07K-207:00 M07K-319:35
    M07K-319:40
Original Abstract: A conjugate compound comprises at least one heat shock
    protein or portion thereof including at least one immunostimulatory
    domain and at least one capsular oligosaccharide or polysaccharide of a
    pathogenic bacteria. The compound comprises oligosaccharides of the
    Meningococci C (MenC) group and a heat shock protein selected from (M.
    bovis) BCG GroEl-type 65kDa hsp (hspR65), recombinant (M. tuberculosis)
    DnaK-type 70kDa hsp (hspR70) and a heat shock protein from (H. pylori).
    The conjugate compounds are useful in the preparation of vaccines to
    prevent bacterial infection.
Claim: A conjugate cpd. comprises at least one heat shock protein (hsp) or
    portion including at least one immunostimulatory domain; and at least
    one oligosaccharide or polysaccharide. The shp may be e.g. M. bovis BCG
    GroEl-type 65 kD hsp (hsp R65), recombinant M. tuberculosis Dnak-type
    60 kD hsp (hsp R70) or a hsp from H. pylori.
Publication No. EP 632727 B1 (Update 199805 E)
Publication Date: 19971229
**KONJUGATE AUS HITZESHOCKPROTEINEN UND OLIGO- ODER POLYSACCHARIDEN
  CONJUGATES FORMED FROM HEAT SHOCK PROTEINS AND OLIGO- OR POLYSACCHARIDES
  COMPOSES CONJUGUES OBTENUS A PARTIR DE PROTEINES DU CHOC THERMIQUE ET
    D'OLIGOSACCHARIDES OU DE POLYSACCHARIDES**
Assignee: BIOCINE S.p.A., Via Fiorentina, 1, I-53100 Siena, IT (BIOC-N)
Inventor: RAPPUOLI, Rino, Via Calamadrei, 39, Quercegrossa, I-53010
   Monteriggioni, IT
  COSTANTINO, Paolo, Via Toscano, 11, I-53034 Colle Val d'Elsa, IT
  VITI, Stefano, Via Dante Alighieri, I-53018 Sovicille, IT
 NORELLI, Francesco, Via Vignali, 16, I-53100 Siena, IT
Agent: Hallybone, Huw George, CARPMAELS AND RANSFORD 43 Bloomsbury Square,
   London WC1A 2RA, GB
Language: EN (27 pages)
Application: EP 1993906489 A 19930308 (Local application)
  WO 1993EP516 A 19930308 (PCT Application)
Priority: IT 1992FI58 A 19920306
Related Publication: WO 1993017712 A (Based on OPI patent )
Designated States: (Regional Original) AT BE CH DE DK ES FR GB GR IE IT LI
    LU MC NL PT SE
Original IPC: A61K-47/48(A) C07K-2/00(B)
Current IPC: A61K-38/00(R,A,N,M,EP,20060101,20051008,A)
    A61K-38/00(R,N,M,EP,20060101,20051008,C)
   A61K-39/00(R,I,M,JP,20060101,20051220,A,L)
   A61K-39/00(R, I, M, JP, 20060101, 20051220, C, L)
    A61K-39/095(R, I, M, EP, 20060101, 20051008, A)
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    A61K-47/48(R, I, M, EP, 20060101, 20051008, A)
    A61K-47/48(R, I, M, EP, 20060101, 20051008, C)
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    A61P-31/04(R, I, M, JP, 20060101, 20051220, A, L)
    A61P-37/00(R,I,M,JP,20060101,20051220,C,L)
    A61P-37/04(R,I,M,JP,20060101,20051220,A,L)
    C07K-14/195(R, I, M, JP, 20060101, 20051220, A, L)
    C07K-14/195(R, I, M, EP, 20060101, 20051008, C)
    C07K-14/205(R, I, M, EP, 20060101, 20051008, A)
    C07K-14/35(R, I, M, EP, 20060101, 20051008, A)
    C07K-14/41(R, I, M, JP, 20060101, 20051220, A, L)
    C07K-14/41(R,I,M,JP,20060101,20051220,C,L)
    C07K-19/00(R, I, M, JP, 20060101, 20051220, A, L)
    C07K-19/00(R, I, M, JP, 20060101, 20051220, C, L)
    C08B-37/00(R, I, M, JP, 20060101, 20051220, A, L)
    C08B-37/00(R,I,M,JP,20060101,20051220,C,L)
    C12N-15/09(R,I,M,JP,20060101,20051220,A,F)
    C12N-15/09(R,I,M,JP,20060101,20051220,C,F)
    C12P-21/02(R,I,M,JP,20060101,20051220,A,L)
    C12P-21/02(R,I,M,JP,20060101,20051220,C,L)
Current ECLA class: A61K-39/095 A61K-47/48R2V C07K-14/205 C07K-14/35
Current ECLA ICO class: K61K-38:00 K61K-39:00 M07K-207:00 M07K-319:35
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- M07K-319:40

 Claim:

 * 1. Konjugatverbindung, die mindestens ein Hitzeschockprotein oder einen

 Teil davon umfasst, enthaltend mindestens eine immunstimulierende

 Dommene und mindestens ein Oligosaccharid oder Polvsaccharid.
 - * 1. A conjugate compound comprising at least one heat shock protein or portion thereof including at least one immunostimulatory domain and at least one oligosaccharide or polysaccharide.

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Publication No. IT 1262896 B (Update 199709 E)
Publication Date: 19960722
Assignee: IST RICERCHE IMMUNOBIOLOGICHE SIENA SRL (RICE-N)
Language: IT
Application: IT 1992FI58 A 19920306 (Local application)
Original IPC: C08G(A)
Current IPC: A61K-38/00(R,A,N,M,EP,20060101,20051008,A)
    A61K-38/00(R,N,M,EP,20060101,20051008,C)
    A61K-39/00(R,I,M,JP,20060101,20051220,A,L)
    A61K-39/00(R,I,M,JP,20060101,20051220,C,L)
    A61K-39/095(R,I,M,EP,20060101,20051008,A)
    A61K-39/095(R, I, M, EP, 20060101, 20051008, C)
    A61K-39/385(R, I, M, JP, 20060101, 20051220, A, L)
    A61K-39/385(R, I, M, JP, 20060101, 20051220, C, L)
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    C07K-14/205(R, I, M, EP, 20060101, 20051008, A)
    C07K-14/35(R, I, M, EP, 20060101, 20051008, A)
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C07K-14/41(R, I, M, JP, 20060101, 20051220, A, L)
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    C12N-15/09(R, I, M, JP, 20060101, 20051220, C, F)
    C12P-21/02(R,I,M,JP,20060101,20051220,A,L)
    C12P-21/02(R,I,M,JP,20060101,20051220,C,L)
Current ECLA class: A61K-39/095 A61K-47/48R2V C07K-14/205 C07K-14/35
Current ECLA ICO class: K61K-38:00 K61K-39:00 M07K-207:00 M07K-319:35
    M07K-319:40
Japan
Publication No. JP 7504423 W (Update 199528 E)
Publication Date: 19950518
Assignee: BIOCINE SCLAVO SPA (ISTS)
Inventor: RAPPUOLI R
 COSTANTINO P
 NORELLI F
Language: JA
Application: JP 1993515333 A 19930308 (Local application)
  WO 1993EP516 A 19930308 (PCT Application)
Priority: IT 1992FI58 A 19920306
Related Publication: WO 1993017712 A (Based on OPI patent )
Original IPC: A61K-47/48(A) A61K-39/385(B) C07K-14/195(B)
Current IPC: A61K-47/48(A) A61K-39/385(B) C07K-14/195(B)
Current ECLA class: A61K-39/095 A61K-47/48R2V C07K-14/205 C07K-14/35
Current ECLA ICO class: K61K-38:00 K61K-39:00 M07K-207:00 M07K-319:35
   M07K-319:40
Publication No. JP 2004346083 A (Update 200481 E)
Publication Date: 20041209
**CONJUGATE FORMED FROM HEAT SHOCK PROTEIN AND OLIGO- OR POLYSACCHARIDE**
Assignee: CHIRON SRL (CHIR-N)
Inventor: RAPPUOLI RINO
  CONSTANTINO PAOLO
  VITI STEFANO
 NORELLI FRANCESCO
Language: JA (36 pages)
Application: JP 1993515333 A 19930308 (Division of application)
  JP 2004216652 A 20040723 (Local application)
Priority: IT 1992FI58 A 19920306
Original IPC: C07K-14/195(A) A61K-39/00(B) C07K-14/35(B) C08B-37/00(B)
    C12P-21/02(B)
Current IPC: A61K-38/00(R,A,N,M,EP,20060101,20051008,A)
    A61K-38/00(R,N,M,EP,20060101,20051008,C)
    A61K-39/00(R,I,M,JP,20060101,20051220,A,L)
    A61K-39/00(R, I, M, JP, 20060101, 20051220, C, L)
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    A61K-39/385(R,I,M,JP,20060101,20051220,A,L)
    A61K-39/385(R,I,M,JP,20060101,20051220,C,L)
    A61K-47/48(R, I, M, EP, 20060101, 20051008, A)
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    A61P-31/04(R,I,M,JP,20060101,20051220,A,L)
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    C07K-14/195(R, I, M, EP, 20060101, 20051008, C)
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C07K-14/205(R, I, M, EP, 20060101, 20051008, A)
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    C07K-19/00(R, I, M, JP, 20060101, 20051220, A, L)
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Current ECLA ICO class: K61K-38:00 K61K-39:00 M07K-207:00 M07K-319:35
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Current JP F-Terms: 4B024 4B064 4C085 4C090 4H045 4B024AA01 4C090AA02
    4C085AA03 4C090AA05 4C090AA09 4H045AA10 4H045AA11 4H045AA20 4H045AA30
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    4B024GA11 4B024GA19 4B024HA03 4B024HA08 4B024HA09 4B024HA14
Publication No. JP 2005068131 A (Update 200520 E)
Publication Date: 20050317
**CONJUGATE FORMED FROM HEAT SHOCK PROTEIN AND OLIGOSACCHARIDE OR
    POLYSACCHARIDE**
Assignee: CHIRON SRL (CHIR-N)
Inventor: RAPPUOLI RINO
  CONSTANTINO PAOLO
  VITI STEFANO
 NORELLI FRANCESCO
Language: JA (35 pages)
Application: JP 1993515333 A 19930308 (Division of application)
  JP 2004137928 A 20040506 (Local application)
Priority: IT 1992FI58 A 19920306
Original IPC: A61K-39/00(A) A61K-47/48(B) A61P-31/04(B) A61P-37/04(B)
    C07K-14/195(B) C07K-14/35(B) C07K-19/00(B)
Current IPC: A61K-38/00(R,A,N,M,EP,20060101,20051008,A)
    A61K-38/00(R,N,M,EP,20060101,20051008,C)
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    A61K-39/385(R,I,M,JP,20060101,20051220,C,L)
    A61K-47/48(R, I, M, EP, 20060101, 20051008, A)
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    C07K-14/195(R,I,M,EP,20060101,20051008,C)
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    C07K-14/35(R, I, M, EP, 20060101, 20051008, A)
    C07K-14/41(R, I, M, JP, 20060101, 20051220, A, L)
    C07K-14/41(R,I,M,JP,20060101,20051220,C,L)
    C07K-19/00(R,I,M,JP,20060101,20051220,A,L)
    C07K-19/00(R,I,M,JP,20060101,20051220,C,L)
    C08B-37/00(R, I, M, JP, 20060101, 20051220, A, L)
    C08B-37/00(R,I,M,JP,20060101,20051220,C,L)
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C12N-15/09(R, I, M, JP, 20060101, 20051220, A, F)
    C12N-15/09(R,I,M,JP,20060101,20051220,C,F)
    C12P-21/02(R, I, M, JP, 20060101, 20051220, A, L)
    C12P-21/02(R,I,M,JP,20060101,20051220,C,L)
Current ECLA class: A61K-39/095 A61K-47/48R2V C07K-14/205 C07K-14/35
Current ECLA ICO class: K61K-38:00 K61K-39:00 M07K-207:00 M07K-319:35
    M07K-319:40
Current JP F-Terms: 4C076 4C085 4C201 4H045 4C085AA03 4H045AA10 4H045AA11
    4C085BA07 4H045BA10 4H045BA41 4C076BB11 4C085BB24 4H045CA11 4C076CC07
    4C085CC21 4C076CC31 4H045DA86 4C085DD51 4H045EA31 4C076EE41 4C076EE59
    4H045FA74 4C076FF70 4C085GG01
Publication No. JP 2009102344 A (Update 200933 E)
Publication Date: 20090514
b CONJUGATE FORMED FROM HEAT SHOCK PROTEIN AND OLIGOSACCHARIDE OR
    POLYSACCHARIDE**
Assignee: CHIRON SRL (CHIR-N)
Inventor: RAPPUOLI RINO
  CONSTANTINO PAOLO
  VITI STEFANO
  NORELLI FRANCESCO
Language: JA (35 pages)
Application: JP 2008312770 A 20081208 (Local application)
  JP 2004137928 A 19930308 (Division of application)
Priority: IT 1992FI58 A 19920306
Original IPC: A61K-39/00(B,I,H,JP,20060101,20090417,A,F)
    A61K-39/00(B, I, M, 98, 20060101, 20090417, C)
    A61P-31/00(B, I, M, 98, 20060101, 20090417, C)
    A61P-31/04(B, I, H, JP, 20060101, 20090417, A, L)
    C07K-14/195(B, N, M, 98, 20060101, 20090417, C)
    C07K-14/35(B, N, H, JP, 20060101, 20090417, A, L)
Current IPC: A61K-38/00(R,N,M,EP,20060101,20051008,A)
    A61K-38/00(R,N,M,EP,20060101,20051008,C)
    A61K-39/00(B, I, H, JP, 20060101, 20090417, A, F)
    A61K-39/00(B, I, H, JP, 20090101, 20090417, C, F)
    A61K-39/095(R, I, M, EP, 20060101, 20051008, A)
    A61K-39/095(R, I, M, EP, 20060101, 20051008, C)
    A61K-39/385(R, I, M, JP, 20060101, 20051220, A, L)
    A61K-39/385(R, I, M, JP, 20060101, 20051220, C, L)
    A61K-47/48(R,I,M,EP,20060101,20051008,A)
    A61K-47/48(R, I, M, EP, 20060101, 20051008, C)
    A61P-31/00(B,I,H,JP,20090101,20090417,C,L)
    A61P-31/04(B, I, H, JP, 20060101, 20090417, A, L)
    A61P-37/00(R, I, M, JP, 20060101, 20051220, C, L)
    A61P-37/04(R, I, M, JP, 20060101, 20051220, A, L)
    C07K-14/195(R, I, M, JP, 20060101, 20051220, A, L)
    C07K-14/195(R,I,M,EP,20060101,20051008,C)
    C07K-14/205(R, I, M, EP, 20060101, 20051008, A)
    C07K-14/35(R, I, M, EP, 20060101, 20051008, A)
    C07K-14/41(R, I, M, JP, 20060101, 20051220, A, L)
    C07K-14/41(R, I, M, JP, 20060101, 20051220, C, L)
    C07K-19/00(R, I, M, JP, 20060101, 20051220, A, L)
    C07K-19/00(R, I, M, JP, 20060101, 20051220, C, L)
    C08B-37/00(R, I, M, JP, 20060101, 20051220, A, L)
    C08B-37/00(R,I,M,JP,20060101,20051220,C,L)
    C12N-15/09(R, I, M, JP, 20060101, 20051220, A, F)
    C12N-15/09(R,I,M,JP,20060101,20051220,C,F)
    C12P-21/02(R,I,M,JP,20060101,20051220,A,L)
    C12P-21/02(R,I,M,JP,20060101,20051220,C,L)
Current ECLA class: A61K-39/095 A61K-47/48R2V C07K-14/205 C07K-14/35
Current ECLA ICO class: K61K-38:00 K61K-39:00 M07K-207:00 M07K-319:35
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M07K-319:40

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Current JP FI-Terms: A61K-39/00 H (main, A, ZNA) A61P-31/04 (secondary, B)
    C07K-14/35 (additional, -)
Current JP F-Terms: 4C085 4C201 4H045 4C085AA03 4C085AA04 4H045AA30
    4H045BA10 4C085BB11 4C085BB24 4H045CA11 4H045EA31 4C085EE03 4H045FA74
Original Abstract: It is achieved by the career carrier used
    conventionally and reliance also identifies the novel protein
    career|carrier which gives a favorable immunogen characteristic
    conjugate, Make possible the reliably highly potent vaccination which
    raises the immunogenic response with respect to an oligosaccharide and
    polysaccharide. The conjugate compound containing some heat-shock
    proteins which contain at least one heat-shock protein or at least one
    immunostimulation domain, and the oligosaccharide or polysaccharide of
    at least one capsule of a pathogenic microbe. Absence (Field|area of
    invention) This invention relates to the conjugate compound which
    consists of a heat-shock protein, and polysaccharide or an
    oligosaccharide (In particular, it is polysaccharide or the
    oligosaccharide derived from a capsule of a pathogenic
    microorganism.).this compound can induce|quide|derive formation of an
    anti- polysaccharide antibody. Therefore, this compound is useful as a
    vaccine used for a human and an animal.
Claim: Invention as described in specification.
Publication No. JP 2011052000 A (Update 201121 E)
Publication Date: 20110317
**Conjugate formed from a heat-shock protein, an oligosaccharide, or
    polysaccharide**
Assignee: CHIRON SPA; JP (CHIR)
Language: JA (35 pages)
Application: JP 2010239145 A 20101025 (Local application)
  JP 2008312770 A 19930308 (Division of application)
Priority: IT 1992FI58 A 19920306
Original IPC: A61K-31/715(B,I,H,JP,20060101,20110218,A,L)
    A61K-39/02(B, I, H, JP, 20060101, 20110218, A, L)
    A61K-47/48(B, I, H, JP, 20060101, 20110218, A, L)
    A61P-31/04(B, I, H, JP, 20060101, 20110218, A, L)
    C07K-14/195(B, I, H, JP, 20060101, 20110218, A, F)
    C12N-15/09(B, I, H, JP, 20060101, 20110218, A, L)
    C12P-19/04(B, N, H, JP, 20060101, 20110218, A, L)
    C12P-21/02(B, N, H, JP, 20060101, 20110218, A, L)
Current IPC: A61K-31/715(B,I,H,JP,20060101,20110218,A,L)
    A61K-39/02(B, I, H, JP, 20060101, 20110218, A, L)
    A61K-47/48(B, I, H, JP, 20060101, 20110218, A, L)
    A61P-31/04(B, I, H, JP, 20060101, 20110218, A, L)
    C07K-14/195(B, I, H, JP, 20060101, 20110218, A, F)
    C12N-15/09(B, I, H, JP, 20060101, 20110218, A, L)
    C12P-19/04(B, N, H, JP, 20060101, 20110218, A, L)
    C12P-21/02(B,N,H,JP,20060101,20110218,A,L)
Current JP FI-Terms: C07K-14/195 (main, A, ZNA) A61K-31/715 (secondary, B)
    A61K-39/02 (secondary, B) A61K-47/48 (secondary, B) A61P-31/04
    (secondary, B) C12N-15/00 A (secondary, B) C12P-19/04 (additional, -)
    C12P-21/02 C (additional, -)
Current JP F-Terms: 4B024 4B064 4C076 4C085 4C086 4H045 4B024AA01 4C086AA01
    4C086AA02 4C085AA03 4C086AA03 4H045AA11 4H045AA20 4H045AA30 4B064AF11
    4B064AG31 4C085BA07 4H045BA10 4B024BA31 4H045BA53 4B024CA02 4B064CA05
    4H045CA11 4B064CA19 4B064CC24 4C076CC31 4B064DA01 4B024DA06 4H045DA86
    4C085DD51 4C085DD62 4C086EA28 4H045EA29 4C085EE01 4C076EE41A 4C076EE59A
    4H045FA10 4H045FA74 4B024GA13 4C085GG03 4C085GG08 4C085GG10 4C086MA02
    4C086MA05 4C086NA14 4C086ZB35
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Original Abstract: It is achieved by the carrier used conventionally and reliance also identifies the novel protein carrier which gives a favorable immunogen characteristic conjugate, Make possible the reliably highly potent vaccination which raises the immunogenic response with

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respect to an oligosaccharide and polysaccharide. The conjugate compound
    containing some heat-shock proteins which contain at least one
    heat-shock protein or at least one immunostimulation domain, and the
    oligosaccharide or polysaccharide of at least one capsule of a
    pathogenic microbe. Absence (Field | area of invention) This invention
    relates to the conjugate compound which consists of a heat-shock
    protein, and polysaccharide or an oligosaccharide (In particular, it is
    polysaccharide or the oligosaccharide derived from a capsule of a
    pathogenic microorganism.).this compound can induce | guide | derive
    formation of an anti- polysaccharide antibody. Therefore, this compound
    is useful as a vaccine used for a human and an animal.
Claim: Invention as described in a specification.
Publication No. JP 3641483 B2 (Update 200527 E)
Publication Date: 20050420
Assignee: CHIRON SPA (CHIR-N)
Language: JA (27 pages)
Application: JP 1993515333 A 19930308 (Local application)
  WO 1993EP516 A 19930308 (PCT Application)
Priority: IT 1992FI58 A 19920306
Related Publication: JP 07504423 A (Previously issued patent)
 WO 1993017712 A (Based on OPI patent )
Original IPC: C08B-37/00(A) A61K-39/385(B) A61K-47/48(B)
Current IPC: A61K-38/00(R.A.N.M.EP.20060101.20051008.A)
    A61K-38/00(R,N,M,EP,20060101,20051008,C)
    A61K-39/00(R,I,M,JP,20060101,20051220,A,L)
    A61K-39/00(R, I, M, JP, 20060101, 20051220, C, L)
    A61K-39/095(R,I,M,EP,20060101,20051008,A)
    A61K-39/095(R,I,M,EP,20060101,20051008,C)
    A61K-39/385(R, I, M, JP, 20060101, 20051220, A, L)
    A61K-39/385(R, I, M, JP, 20060101, 20051220, C, L)
    A61K-47/48(R, I, M, EP, 20060101, 20051008, A)
   A61K-47/48(R,I,M,EP,20060101,20051008,C)
    A61P-31/00(R,I,M,JP,20060101,20051220,C,L)
    A61P-31/04(R, I, M, JP, 20060101, 20051220, A, L)
    A61P-37/00(R, I, M, JP, 20060101, 20051220, C, L)
   A61P-37/04(R, I, M, JP, 20060101, 20051220, A, L)
    C07K-14/195(R, I, M, JP, 20060101, 20051220, A, L)
    C07K-14/195(R, I, M, EP, 20060101, 20051008, C)
    C07K-14/205(R, I, M, EP, 20060101, 20051008, A)
    C07K-14/35(R,I,M,EP,20060101,20051008,A)
    C07K-14/41(R, I, M, JP, 20060101, 20051220, A, L)
    C07K-14/41(R, I, M, JP, 20060101, 20051220, C, L)
    C07K-19/00(R, I, M, JP, 20060101, 20051220, A, L)
    C07K-19/00(R, I, M, JP, 20060101, 20051220, C, L)
    C08B-37/00(R,I,M,JP,20060101,20051220,A,L)
    C08B-37/00(R,I,M,JP,20060101,20051220,C,L)
    C12N-15/09(R,I,M,JP,20060101,20051220,A,F)
    C12N-15/09(R, I, M, JP, 20060101, 20051220, C, F)
    C12P-21/02(R, I, M, JP, 20060101, 20051220, A, L)
    C12P-21/02(R, I, M, JP, 20060101, 20051220, C, L)
Current ECLA class: A61K-39/095 A61K-47/48R2V C07K-14/205 C07K-14/35
Current ECLA ICO class: K61K-38:00 K61K-39:00 M07K-207:00 M07K-319:35
    M07K-319:40
Current JP F-Terms: 4C076 4C085 4C090 4H045 4C090AA02 4C085AA03 4C085AA04
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    4C090BB65 4C090BB69 4C090BC19 4C090BC20 4H045CA11 4C090CA35 4C085CC05
    4C076CC06 4C085CC07 4C085CC08 4C085CC21 4C085CC32 4C085CC33 4C090DA23
    4C085DD51 4C085DD52 4C085DD59 4C085DD62 4C085DD86 4H045EA29 4H045EA31
    4C076EE41A 4C076EE59A 4H045FA74 4C085FF01 4C085FF02 4C085FF03 4C085FF12
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United States
Publication No. US 6403099 B1 (Update 200244 E)
Publication Date: 20020611
**Conjugates formed from heat shock proteins and oligo-or
    polysaccharides. **
Assignee: Chiron S.p.A., Siena, IT (CHIR)
Inventor: Rappuoli, Rino, Ouercegrossa, IT
  Costantino, Paolo, Colle dorimeElsa, IT
 Viti, Stefano, Sovicille, IT
 Norelli, Francesco, Siena, IT
Agent: Attwell; Gwilym J.O.
  Harbin; Alisa A.
  Blackburn: Robert P.
Language: EN
Application: WO 1993EP516 A 19930308 (PCT Application)
  US 1994256847 A 19941101 (Local application)
Priority: IT 1992FI58 A 19920306
Related Publication: WO 1993017712 A (Based on OPI patent )
Original IPC: C01B-3/00(A) C07K-1/00(B) G01N-33/554(B)
Current IPC: A61K-38/00(R,A,N,M,EP,20060101,20051008,A)
    A61K-38/00(R,N,M,EP,20060101,20051008,C)
    A61K-39/00(R, I, M, JP, 20060101, 20051220, A, L)
    A61K-39/00(R, I, M, JP, 20060101, 20051220, C, L)
    A61K-39/095(R, I, M, EP, 20060101, 20051008, A)
    A61K-39/095(R,I,M,EP,20060101,20051008,C)
    A61K-39/385(R,I,M,JP,20060101,20051220,A,L)
    A61K-39/385(R, I, M, JP, 20060101, 20051220, C, L)
    A61K-47/48(R,I,M,EP,20060101,20051008,A)
    A61K-47/48(R, I, M, EP, 20060101, 20051008, C)
    A61P-31/00(R, I, M, JP, 20060101, 20051220, C, L)
    A61P-31/04(R,I,M,JP,20060101,20051220,A,L)
    A61P-37/00(R, I, M, JP, 20060101, 20051220, C, L)
    A61P-37/04(R, I, M, JP, 20060101, 20051220, A, L)
    C07K-14/195(R, I, M, JP, 20060101, 20051220, A, L)
    C07K-14/195(R, I, M, EP, 20060101, 20051008, C)
    C07K-14/205(R, I, M, EP, 20060101, 20051008, A)
    C07K-14/35(R, I, M, EP, 20060101, 20051008, A)
    C07K-14/41(R, I, M, JP, 20060101, 20051220, A, L)
    C07K-14/41(R,I,M,JP,20060101,20051220,C,L)
    C07K-19/00(R, I, M, JP, 20060101, 20051220, A, L)
    C07K-19/00(R, I, M, JP, 20060101, 20051220, C, L)
    C08B-37/00(R, I, M, JP, 20060101, 20051220, A, L)
    C08B-37/00(R, I, M, JP, 20060101, 20051220, C, L)
    C12N-15/09(R, I, M, JP, 20060101, 20051220, A, F)
    C12N-15/09(R, I, M, JP, 20060101, 20051220, C, F)
    C12P-21/02(R, I, M, JP, 20060101, 20051220, A, L)
    C12P-21/02(R, I, M, JP, 20060101, 20051220, C, L)
Current ECLA class: A61K-39/095 A61K-47/48R2V C07K-14/205 C07K-14/35
Current ECLA ICO class: K61K-38:00 K61K-39:00 M07K-207:00 M07K-319:35
    M07K-319:40
Original US Class (main): 424248.1
Original US Class (secondary): 424192.1 530395 514569 4357.32 43512
Original Abstract: The present invention provides conjugate compounds
    comprising at least one heat shock protein or portion thereof including
    at least one immunostimulatory domain and at least one capsular
    oligosaccharide or polysaccharide of a pathogenic bacteria. The
    compound comprises oligosaccharides of the Meningococci C (MenC) group
    and a heat shock protein selected from ~M. bovis ~BCG GroE1-type 65 kDa
    hsp (hspR65), recombinant ~M. tuberculosis ~DnaK-type 70 kDa hsp
    (hspR70) and a heat shock protein from ~H. pylori~. The invention also
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provides processes for producing conjugate compounds, pharmaceutical compositions comprising conjugate compounds, therapeutic compositions comprising conjugate compounds, and methods of inducing an immune response.

Claim:

1.A conjugate compound comprising a portion of at least 11 to 15 amino acid residues of a heat shock protein selected from the group consisting of

~M. bovis ~BCG GroEL-type 65 kDa heat shock protein

and recombinant ~M. tuberculosis ~DnaK-type 70 kDa heat shock protein, wherein said heat shock protein portion includes at least one immunostimulatory domain, said conjugate compound also comprising at least one capsular oligosaccharide or capsular polysaccharide, or immunogenic portion thereof.

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Publication No. WO 1993017712 A2 (Update 199338 B)
Publication Date: 19930916
**CONJUGATES FORMED FROM HEAT SHOCK PROTEINS AND OLIGO- OR
    POLYSACCHARIDES**
Assignee: BIOCINE SCLAVO SPA, IT (ISTS)
Inventor: RAPPUOLI, RINO, IT
  COSTANTINO, PAOLO, IT
  NORELLI, FRANCESCO, IT
  VITI, STEFANO, IT
Language: EN (69 pages, 0 drawings)
Application: WO 1993EP516 A 19930308 (Local application)
Priority: IT 1992FI58 A 19920306
Designated States: (National Original) AT AU BB BG BR CA CH CZ DE DK ES FI
    GB HU JP KP KR LK LU MG MN MW NL NO NZ PL PT RO RU SD SE SK UA US
  (Regional Original) AT BE CH DE DK ES FR GB GR IE IT LU MC NL OA PT SE
Original IPC: A61K-47/48(A) C07K-15/00(B)
Current IPC: A61K-38/00(R,A,N,M,EP,20060101,20051008,A)
    A61K-38/00(R,N,M,EP,20060101,20051008,C)
    A61K-39/00(R, I, M, JP, 20060101, 20051220, A, L)
    A61K-39/00(R, I, M, JP, 20060101, 20051220, C, L)
    A61K-39/095(R, I, M, EP, 20060101, 20051008, A)
    A61K-39/095(R, I, M, EP, 20060101, 20051008, C)
    A61K-39/385(R, I, M, JP, 20060101, 20051220, A, L)
    A61K-39/385(R, I, M, JP, 20060101, 20051220, C, L)
    A61K-47/48(R, I, M, EP, 20060101, 20051008, A)
    A61K-47/48(R,I,M,EP,20060101,20051008,C)
    A61P-31/00(R, I, M, JP, 20060101, 20051220, C, L)
    A61P-31/04(R, I, M, JP, 20060101, 20051220, A, L)
    A61P-37/00(R, I, M, JP, 20060101, 20051220, C, L)
    A61P-37/04(R,I,M,JP,20060101,20051220,A,L)
    C07K-14/195(R, I, M, JP, 20060101, 20051220, A, L)
    C07K-14/195(R, I, M, EP, 20060101, 20051008, C)
    C07K-14/205(R, I, M, EP, 20060101, 20051008, A)
    C07K-14/35(R, I, M, EP, 20060101, 20051008, A)
    C07K-14/41(R, I, M, JP, 20060101, 20051220, A, L)
    C07K-14/41(R, I, M, JP, 20060101, 20051220, C, L)
    C07K-19/00(R, I, M, JP, 20060101, 20051220, A, L)
    C07K-19/00(R, I, M, JP, 20060101, 20051220, C, L)
    C08B-37/00(R,I,M,JP,20060101,20051220,A,L)
    C08B-37/00(R, I, M, JP, 20060101, 20051220, C, L)
    C12N-15/09(R,I,M,JP,20060101,20051220,A,F)
    C12N-15/09(R,I,M,JP,20060101,20051220,C,F)
    C12P-21/02(R,I,M,JP,20060101,20051220,A,L)
    C12P-21/02(R,I,M,JP,20060101,20051220,C,L)
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Current ECLA class: A61K-39/095 A61K-47/48R2V C07K-14/205 C07K-14/35 Current ECLA ICO class: K61K-38:00 K61K-39:00 M07K-207:00 M07K-319:35

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M07K-319:40
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Original Abstract: A conjugate compound comprises at least one heat shock protein or portion thereof including at least one immunostimulatory domain and at least one capsular oligosaccharide or polysaccharide of a pathogenic bacteria. The compound comprises oligosaccharides of the Meningococci C (MenC) group and a heat shock protein selected from (M. bovis) BCG GroEl-type 65kDa hsp (hspR65), recombinant (M. tuberculosis) DaAK-type 70kDa hsp (hspR70) and a heat shock protein from (H. pylori). The conjugate compounds are useful in the preparation of vaccines to

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prevent bacterial infection.
Publication No. WO 1993017712 A3 (Update 199514 E)
Publication Date: 19931111
Assignee: BIOCINE SCLAVO SPA (ISTS)
Inventor: RAPPUOLI R
  COSTANTINO P
 NORELLT F
Language: EN
Application: WO 1993EP516 A 19930308 (Local application)
Priority: IT 1992FI58 A 19920306
Original IPC: A61K-47/48(A) C07K-15/00(B)
Current IPC: A61K-38/00(R,A,N,M,EP,20060101,20051008,A)
    A61K-38/00(R,N,M,EP,20060101,20051008,C)
    A61K-39/00(R, I, M, JP, 20060101, 20051220, A, L)
    A61K-39/00(R, I, M, JP, 20060101, 20051220, C, L)
    A61K-39/095(R, I, M, EP, 20060101, 20051008, A)
    A61K-39/095(R, I, M, EP, 20060101, 20051008, C)
    A61K-39/385(R,I,M,JP,20060101,20051220,A,L)
    A61K-39/385(R, I, M, JP, 20060101, 20051220, C, L)
    A61K-47/48(R,I,M,EP,20060101,20051008,A)
    A61K-47/48(R, I, M, EP, 20060101, 20051008, C)
    A61P-31/00(R, I, M, JP, 20060101, 20051220, C, L)
    A61P-31/04(R,I,M,JP,20060101,20051220,A,L)
    A61P-37/00(R, I, M, JP, 20060101, 20051220, C, L)
    A61P-37/04(R, I, M, JP, 20060101, 20051220, A, L)
    C07K-14/195(R, I, M, JP, 20060101, 20051220, A, L)
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    C07K-14/205(R, I, M, EP, 20060101, 20051008, A)
    C07K-14/35(R, I, M, EP, 20060101, 20051008, A)
    C07K-14/41(R,I,M,JP,20060101,20051220,A,L)
    C07K-14/41(R,I,M,JP,20060101,20051220,C,L)
    C07K-19/00(R, I, M, JP, 20060101, 20051220, A, L)
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    C08B-37/00(R, I, M, JP, 20060101, 20051220, A, L)
    C08B-37/00(R,I,M,JP,20060101,20051220,C,L)
    C12N-15/09(R, I, M, JP, 20060101, 20051220, A, F)
    C12N-15/09(R,I,M,JP,20060101,20051220,C,F)
    C12P-21/02(R, I, M, JP, 20060101, 20051220, A, L)
    C12P-21/02(R, I, M, JP, 20060101, 20051220, C, L)
Current ECLA class: A61K-39/095 A61K-47/48R2V C07K-14/205 C07K-14/35
Current ECLA ICO class: K61K-38:00 K61K-39:00 M07K-207:00 M07K-319:35
    M07K-319:40
 10/7/17
              (Item 17 from file: 351)
DIALOG(R)File 351:Derwent WPI
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0003920445
WPI ACC NO: 1987-009027/198702
XRAM Acc No: C1987-003413
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New glycoproteinic conjugates - prepd. from protein antigen and

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oligosaccharide hapten(s), derived from ****capsular****
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****polysaccharide**** of Gram-positive and Gram-negative bacteria Patent Assignee: IST SIEROTERAPBUTICO & VACCINOGENO (ISTS) Inventor: COSTANTINO P: PORRO M

Patent Family (8 patents, 13 countries)

Patent Paking (o patents, 13 countries)

Application

Pat	ent			App	plication				
Number		Kind	Date	Number		Kind	Date	Update	
EΡ	208375	A	19870114	EP	1986201160	A	19860702	198702	В
JP	62030726	A	19870209	JP	1986156342	A	19860704	198711	E
US	4711779	A	19871208	US	1986881091	A	19860702	198751	E
CA	1272952	A	19900821					199039	E
IΤ	1187753	В	19871223	IT	198521451	A	19850705	199044	E
EP	208375	В	19911211	EP	1986201160	A	19860702	199150	E
DE	3682838	G	19920123					199205	E
JΡ	1995121870	B2	19951225	JP	1986156342	A	19860704	199605	E

Priority Applications (number, kind, date): IT 198521451 A 19850705

Patent Details

Number Kind Lan Pg Dwg Filing Notes

EP 208375 A EN 12 4
Regional Designated States, Original: AT BE CH DE FR GB LI LU NL SE

US 4711779 A EN CA 1272952 A EN

EP 208375 B EN Regional Designated States, Original: AT BE CH DE FR GB LI LU NL SE

Regional Designated States,Original: AT BE CH DE FR GB LI LU NL SE JP 1995121870 B2 JA 9 0 Based on OPI patent JP 62030726

Alerting Abstract EP A

New glycoproteinic conjugates (I), with trivalent immunogenic activity are obtd. by covalent binding of a proteinic antigen, namely CRM 197, tetanus toxoid or pertussis toxin, with an oligosaccharidic hapten derived from the ****capsular**** ****polysaccharide**** of a Gram-positive bacterium and with one derived from the ****capsular****

****polysaccharide**** of a Gram-negative bacterium. The haptens are first activated by introduction of terminal ester gps.

Pref. Gram-positive bacteria are Streptococcus pneumoniae and S. beta-emoliticus. Pref. Gram-negative bacteria are ****Nesiseria**** ****meningitidis****, Haemophilus influenzae, Pseudomonas aeruginosa and E.coli.

USE/ADVANTAGE - (I) are useful as vaccines against capsulate Gram-positive and Gram-negative bacteria, partic. meningococcus and pneumococcus.

Equivalent Alerting Abstract US A

Glycoprotein conjugates are obtd. by linking an antigenic protein (e.g. CRM 197, tetanus toxoid or pertussis toxin) covalently with one or more oligosaccharidic haptene (obtd. from Gram positive bacterial ****capsular**** ****polysaccharide****) and at least one other

oligosaccharidic haptene (from the ****capsular**** ****polysaccharide**** of a Gram negative microorganisms). Both oligosaccharide haptenes are previously activated by introdn. of terminal ester gps.

USE - The prods. are components for triple immunisation vaccines. (9pp)o

Title Terms/Index Terms/Additional Words: NEW; CONJUGATE; PREPARATION; PROTEIN; ANTIGEN; OLIGOSACCHARIDE; HAPTEN; DERIVATIVE; CAPSULE; ****POLYSACCHARIDE****; GRAM; POSITIVE; NEGATIVE; BACTERIA

Class Codes

International Classification (Main): A61K-039/116

(Additional/Secondary): A61K-039/02, A61K-039/05, A61K-039/08, A61K-039/10 International Classification (+ Attributes)

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IPC + Level Value Position Status Version
 A61K-0039/116 A I R 20060101
 C07K-0001/08 A I L R 20060101
 C07K-0001/107 A I L R 20060101
 C07K-0001/113 A I L R 20060101
 C07K-0014/00 A I F R 20060101
 C07K-0014/195 A I L R 20060101
  C07K-0014/41 A I L R 20060101
  C07K-0019/00 A I L R 20060101
 A61K-0039/116 C I
                        R 20060101
 C07K-0001/00 C I L R 20060101
  C07K-0014/00 C I F R 20060101
 C07K-0014/195 C I L R 20060101
 C07K-0014/41 C I L R 20060101
 C07K-0019/00 C I L R 20060101
ECLA: A61K-039/116
US Classification, Issued: 42492, 530395, 530397, 530406
JP Classification
 FI Term
                   Facet Rank Type
A61K-039/02
                          Z indexing
                          Z indexing
A61K-039/05
                          Z indexing
A61K-039/08
                          Z indexing
A61K-039/09
                         Z indexing
Z indexing
Z indexing
Z indexing
Z indexing
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A61K-039/104
A61K-039/108
A61K-039/116
C07K-001/08
C07K-001/107
C07K-001/113
C07K-014/00
C07K-014/195
C07K-014/41
C07K-019/00
F-Term View Point Additional
Theme
       + Figure Code
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          EA31
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 4H045
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File Segment: CPI
DWPI Class: B04; D16
Manual Codes (CPI/A-M): B02-V02; B04-B04C1; B04-C02F; B12-A01; D05-H07
Original Publication Data by Authority
Canada
Publication No. CA 1272952 A (Update 199039 E)
Publication Date: 19900821
Language: EN
Priority: IT 198521451 A 19850705
Current IPC: A61K-39/116(R, I, M, EP, 20060101, 20051008, A)
    A61K-39/116(R, I, M, EP, 20060101, 20051008, C)
    C07K-1/00(R, I, M, JP, 20060101, 20051220, C, L)
    C07K-1/08(R, I, M, JP, 20060101, 20051220, A, L)
    C07K-1/107(R, I, M, JP, 20060101, 20051220, A, L)
    C07K-1/113(R, I, M, JP, 20060101, 20051220, A, L)
    C07K-14/00(R, I, M, JP, 20060101, 20051220, A, F)
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    C07K-14/195(R, I, M, JP, 20060101, 20051220, A, L)
    C07K-14/195(R, I, M, JP, 20060101, 20051220, C, L)
    C07K-14/41(R, I, M, JP, 20060101, 20051220, A, L)
    C07K-14/41(R, I, M, JP, 20060101, 20051220, C, L)
    C07K-19/00(R, I, M, JP, 20060101, 20051220, A, L)
    C07K-19/00(R, I, M, JP, 20060101, 20051220, C, L)
Current ECLA class: A61K-39/116
Germany
Publication No. DE 3682838 G (Update 199205 E)
Publication Date: 19920123
Language: DE
Priority: IT 198521451 A 19850705
Publication No. EP 208375 A (Update 198702 B)
Publication Date: 19870114
**Glycoproteinkonjugate mit trivalenter immunogener Aktivitaet
 Glycoproteinic conjugates having trivalent immunogenic activity
  Conjugues glycoproteiniques avant une activite immunogene trivalente **
Assignee: IST SIEROTERAPEUTICO VACCINOGENO (ISTS)
  SCLAVO S.p.A., Via Fiorentina 1, I-53100 Siena, IT
Inventor: Porro, Massimo, Via S. Marco 43, I-56010 Localita' Collanza
    Asciano Siena, IT
  Costantino, Paolo, Via Toscana 11, I-53034 Colle Di Val D'Elsa Siena, IT
Agent: Roggero, Sergio, et al, Ing. Barzano Zanardo Milano S.p.A. Via
    Borgonuovo 10, I-20121 Milano, IT
Language: EN (12 pages, 4 drawings)
Application: EP 1986201160 A 19860702 (Local application)
Priority: IT 198521451 A 19850705
Designated States: (Regional Original) AT BE CH DE FR GB LI LU NL SE
Original IPC: A61K-39/11 C07K-3/08 C07K-15/14 C07K-17/10 C12P-0/00
Current IPC: A61K-39/116(R,I,M,EP,20060101,20051008,A)
    A61K-39/116(R, I, M, EP, 20060101, 20051008, C)
    C07K-1/00(R, I, M, JP, 20060101, 20051220, C, L)
    C07K-1/08(R, I, M, JP, 20060101, 20051220, A, L)
    C07K-1/107(R,I,M,JP,20060101,20051220,A,L)
    C07K-1/113(R,I,M,JP,20060101,20051220,A,L)
    C07K-14/00(R,I,M,JP,20060101,20051220,A,F)
    C07K-14/00(R, I, M, JP, 20060101, 20051220, C, F)
    C07K-14/195(R, I, M, JP, 20060101, 20051220, A, L)
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C07K-14/195(R,I,M,JP,20060101,20051220,C,L)
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    C07K-14/41(R,I,M,JP,20060101,20051220,C,L)
    C07K-19/00(R,I,M,JP,20060101,20051220,A,L)
    C07K-19/00(R,I,M,JP,20060101,20051220,C,L)
Current ECLA class: A61K-39/116
Original Abstract: Glycoproteinic conjugates having trivalent immunogenic
    activity obtained by binding, by a covalent bond, to a protein selected
    among CRM 197, tetanus toxoid, and pertussis toxin, at least an
    oligosaccharidic hapten derived from the capsular polysaccharide of a
    gram-positive bacterium and at least an oligosaccharidic hapten derived
    from the capsular polysaccharide of a gram-negative bacterium, and
    wherein said oligosaccharidic haptens are previously activated by
    introducing terminal esters.
Claim: New glycoproteinic conjugates (I), with trivalent immunogenic
    activity are obtd. by covalent binding of a proteinic antigen, namely
    CRM 197, tetanus toxoid or pertussis toxin, with an oligosaccharidic
    hapten derived from the capsular polysaccharide of a Gram-positive
    bacterium and with one derived from the capsular polysaccharide of a
    Gram-negative bacterium. The haptens are first activated by
    introduction of terminal ester qps.
  Pref. Gram-positive bacteria are Streptococcus pneumoniae and S.
    beta-emoliticus. Pref. Gram-negative bacteria are Neisseria
    meningitidis, Haemophilus influenzae, Pseudomonas aeruginosa and
    E.coli.
Publication No. EP 208375 B (Update 199150 E)
Publication Date: 19911211
**Glycoproteinkonjugate mit trivalenter immunogener Aktivitaet
  Glycoproteinic conjugates having trivalent immunogenic activity
  Conjugues glycoproteiniques ayant une activite immunogene trivalente **
Assignee: SCLAVO S.p.A., Via Fiorentina 1, I-53100 Siena, IT
Inventor: Porro, Massimo, Via S. Marco 43, I-56010 Localita' Collanza
    Asciano Siena, IT
  Costantino, Paolo, Via Toscana 11, I-53034 Colle Di Val D'Elsa Siena, IT
Agent: Gervasi, Gemma, Dr. et al, Studio Brevetti e Marchi NOTARBARTOLO
   GERVASI 33, Viale Bianca Maria, I-20122 Milano, IT
Language: EN
Application: EP 1986201160 A 19860702 (Local application)
Priority: IT 198521451 A 19850705
Designated States: (Regional Original) AT BE CH DE FR GB LI LU NL SE
Original IPC: A61K-39/116 A61K-39/385 C07K-15/14
Current IPC: A61K-39/116(R,A,I,M,EP,20060101,20051008,A)
   A61K-39/116(R, I, M, EP, 20060101, 20051008, C)
    C07K-1/00(R,I,M,JP,20060101,20051220,C,L)
    C07K-1/08(R,I,M,JP,20060101,20051220,A,L)
    C07K-1/107(R,I,M,JP,20060101,20051220,A,L)
    C07K-1/113(R,I,M,JP,20060101,20051220,A,L)
    C07K-14/00(R, I, M, JP, 20060101, 20051220, A, F)
    C07K-14/00(R, I, M, JP, 20060101, 20051220, C, F)
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    20060101,20051220,C,L) C07K-14/41(R,I,M,JP,20060101,20051220,A,L)
    C07K-14/41(R, I, M, JP, 20060101, 20051220, C, L)
    C07K-19/00(R, I, M, JP, 20060101, 20051220, A, L)
    C07K-19/00(R, I, M, JP, 20060101, 20051220, C, L)
Current ECLA class: A61K-39/116
```

1. Glycoproteinkonjugate mit trivalenter Immunogenaktivitaet, erhalten durch Binden wenigatens eines Oligosaccharidhaptens, stammend vom kapsulaeren Polysaccharid eines gram-positiven Bakteriums, und wenigstens eines Oligosaccharidhaptens, stammend vom kapsulaeren Polysaccharid eines gram-negativen Bakteriums, durch eine kovalente

Claim:

Bindung an ein Proteinantigen, ausgewaehlt aus CBM 197, Tetanustoxoid oder Pertussistoxin, nach der Voraktivierung der Oligosaccharidhaptene durch Einfuehren von endstaendigen Estergruppen.

* 1. Glycoproteinic conjugates having trivalent immunogenic activity, obtained by binding, by a covalent bond, to a proteinic antigen selected among CRM 197, tetanus toxoid, or pertussis toxin, at least one oligosaccharidic hapten derived from the capsular polysaccharide of a gram-positive bacterium, and at least one oligosaccharidic hapten derived from the capsular polysaccharide of a gram-negative bacterium, after the preliminary activation of the said oligosaccharidic haptens by the introduction of terminal ester groups.

```
Publication No. IT 1187753 B (Update 199044 E)
Publication Date: 19871223
Language: IT
Application: IT 198521451 A 19850705
Priority: IT 198521451 A 19850705
Current IPC: A61K-39/116(R.I.M.EP.20060101.20051008.A)
    A61K-39/116(R, I, M, EP, 20060101, 20051008, C)
    C07K-1/00(R, I, M, JP, 20060101, 20051220, C, L)
    C07K-1/08(R, I, M, JP, 20060101, 20051220, A, L)
    C07K-1/107(R, I, M, JP, 20060101, 20051220, A, L)
    C07K-1/113(R, I, M, JP, 20060101, 20051220, A, L)
    C07K-14/00(R, I, M, JP, 20060101, 20051220, A, F)
    C07K-14/00(R, I, M, JP, 20060101, 20051220, C, F)
    C07K-14/195(R, I, M, JP, 20060101, 20051220, A, L)
    C07K-14/195(R, I, M, JP, 20060101, 20051220, C, L)
    C07K-14/41(R,I,M,JP,20060101,20051220,A,L)
    C07K-14/41(R,I,M,JP,20060101,20051220,C,L)
    C07K-19/00(R, I, M, JP, 20060101, 20051220, A, L)
    C07K-19/00(R, I, M, JP, 20060101, 20051220, C, L)
Current ECLA class: A61K-39/116
Japan
Publication No. JP 62030726 A (Update 198711 E)
Publication Date: 19870209
Language: JA
Application: JP 1986156342 A 19860704 (Local application)
Priority: IT 198521451 A 19850705
Current IPC: A61K-39/116(R, I, M, EP, 20060101, 20051008, A)
    A61K-39/116(R, I, M, EP, 20060101, 20051008, C)
    C07K-1/00(R,I,M,JP,20060101,20051220,C,L)
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    C07K-1/107(R, I, M, JP, 20060101, 20051220, A, L)
    C07K-1/113(R, I, M, JP, 20060101, 20051220, A, L)
    C07K-14/00(R, I, M, JP, 20060101, 20051220, A, F)
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    C07K-19/00(R,I,M,JP,20060101,20051220,A,L)
    C07K-19/00(R,I,M,JP,20060101,20051220,C,L)
Current ECLA class: A61K-39/116
Current JP FI-Terms: A61K-39:02 (indexing, Z) A61K-39:05 (indexing, Z)
    A61K-39:08 (indexing, Z) A61K-39:09 (indexing, Z) A61K-39:095
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(indexing, Z) A61K-39:10 (indexing, Z) A61K-39:104 (indexing, Z)
    A61K-39:108 (indexing, Z) A61K-39/116 C07K-1/08 C07K-1/107 C07K-1/113
    C07K-14/00 C07K-14/195 C07K-14/41 C07K-19/00
Current JP F-Terms: 4C085 4H045 4C085AA04 4H045AA10 4H045AA30 4C085BA13
    4C085BA14 4C085BA18 4C085BA21 4H045BA41 4H045BA53 4H045CA11 4C085CC07
    4H045DA86 4C085DD03 4C085DD18 4H045EA31 4C085EE03 4H045FA20 4H045FA41
    4H045FA50 4C085GG06
Publication No. JP 1995121870 B2 (Update 199605 E)
Publication Date: 19951225
Assignee: IST SIEROTERAPEUTICO & VACCINOGENO (ISTS)
Inventor: PORRO M
 COSTANTINO P
Language: JA (9 pages, 0 drawings)
Application: JP 1986156342 A 19860704 (Local application)
Priority: IT 198521451 A 19850705
Related Publication: JP 62030726 A (Based on OPI patent )
Original IPC: A61K-39/116(A) C07K-1/08(B) C07K-14/195(B) A61K-39/116(C)
    A61K-39:02(C) A61K-39:05(C) A61K-39/116(D) A61K-39:02(D) A61K-39:08(D)
    A61K-39/116(E) A61K-39:02(E) A61K-39:10(E)
Current IPC: A61K-39/116(A) C07K-1/08(B) C07K-14/195(B) A61K-39/116(C)
    A61K-39:02(C) A61K-39:05(C) A61K-39/116(D) A61K-39:02(D) A61K-39:08(D)
    A61K-39/116(E) A61K-39:02(E) A61K-39:10(E)
United States
Publication No. US 4711779 A (Update 198751 E)
Publication Date: 19871208
**Glycoproteinic conjugates having trivalent immunogenic activity**
Assignee: Sclavo S.p.A.
Inventor: Porro, Massimo, IT
 Costantino, Paolo
Agent: Hedman, Gibson, Costigan Hoare
Language: EN (9 pages)
Application: US 1986881091 A 19860702 (Local application)
Priority: IT 198521451 A 19850705
Original IPC: C07K-17/10 A61K-39/385
Current IPC: A61K-39/116(R,A,I,M,EP,20060101,20051008,A)
    A61K-39/116(R, I, M, EP, 20060101, 20051008, C)
    C07K-1/00(R,I,M,JP,20060101,20051220,C,L)
    C07K-1/08(R,I,M,JP,20060101,20051220,A,L)
    C07K-1/107(R,I,M,JP,20060101,20051220,A,L)
   C07K-1/113(R, I, M, JP, 20060101, 20051220, A, L)
    C07K-14/00(R, I, M, JP, 20060101, 20051220, A, F)
    C07K-14/00(R,I,M,JP,20060101,20051220,C,F)
    C07K-14/195(R,I,M,JP,20060101,20051220,A,L)
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    C07K-14/41(R,I,M,JP,20060101,20051220,A,L)
    C07K-14/41(R, I, M, JP, 20060101, 20051220, C, L)
    C07K-19/00(R, I, M, JP, 20060101, 20051220, A, L)
    C07K-19/00(R, I, M, JP, 20060101, 20051220, C, L)
Current ECLA class: A61K-39/116
Original US Class (main): 42492
Original US Class (secondary): 530395 530397 530406
Original Abstract: Glycoproteinic conjugates having trivalent immunogenic
    activity obtained by binding, by a covalent bond, to a protein selected
    among CRM 197, tetanus toxoid, and pertussis toxin, at least an
    oligosaccharidic hapten derived from the capsular polysaccharide of a
    gram-positive bacterium and at least an oligosaccharidic hapten derived
    from the capsular polysaccharide of a gram-negative bacterium, and
   wherein said oligosaccharidic haptens are previously activated by
    introducing terminal esters.
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10/7/18 (Item 1 from file: 24) DIALOG(R)File 24:CSA Life Sciences Abstracts (c) 2011 CSA. All rts. reserv.

0004004471 IP ACCESSION NO: 11359805

Toll-like receptor 2 dependent immunogenicity of glycoconjugate vaccines containing chemically derived zwitterionic polysaccharides

Gallorini, Simona; ****Berti, Francesco****; Mancuso, Giuseppe; Cozzi, Roberta; Tortoli, Marco; Volpini, Gianfranco; Telford, John L; Beninati, Concetta; Maione, Domenico; Wack, Andreas Novartis Vaccines Research Center, Via Fiorentina 1, 53100 Siena, Italy, [mailto:awackenim: mc.ac.uk]

Proceedings of the National Academy of Sciences, USA, v 106, n 41, p 17481-17486, January , 2009 PUBLICATION DATE: 2009

PUBLISHER: National Academy of Sciences, 2101 Constitution Ave. Washington DC 20418 USA

DOCUMENT TYPE: Journal Article RECORD TYPE: Abstract LANGUAGE: English SUMMARY LANGUAGE: English ISSN: 0027-8424 DOI: 10.1073/pnas.0903313106

FILE SEGMENT: Bacteriology Abstracts (Microbiology B); Immunology Abstracts

ABSTRACT:

Group B Streptococcus (GBS) causes serious infection in neonates and is an important target of vaccine development. Zwitterionic polysaccharides (ZPS), obtained through chemical introduction of positive charges into anionic polysaccharides (PS) from GBS, have the abilityto activate human and mouse antigen presenting cells (APCs) through toll-like receptor 2 (TLR2). To generate a polysaccharidevaccine with antigen (Ag) and adjuvant properties in one molecule, we have conjugated ZPS with a carrier protein. ZPS-glycoconjugatesinduce higher T-cell and Ab responses to carrier and PS, respectively, compared to control PS-glycoconjugates made with thenative ****polysaccharide**** form. The increased immunogenicity of ZPS-conjugates correlates with their ability to activate dendriticcells (DCs). Moreover, protection of mothers or neonate offspring from lethal GBS challenge is better when mothers are immunized with ZPS-conjugates compared to immunization with PS-conjugates. In TLR2 knockout mice, ZPS-conjugates lose both their increasedimmunogenicity and protective effect after vaccination. When ZPS are coadministered as adjuvants with unconjugated tetanustoxoid (TT), they have the ability to increase the TT-specific antibody titer. In conclusion, glycoconjugates containing ZPSare potent vaccines. They target Ag to TLR2-expressing APCs and activate these APCs, leading to better T-cell priming andultimately to higher protective Ab titers. Thus, rational chemical design can generate potent PS-adjuvants with wide application, including glycoconjugates and coadministration with unrelated protein Ags.

10/7/19 (Item 2 from file: 24) DIALOG(R)File 24:CSA Life Sciences Abstracts (c) 2011 CSA. All rts. reserv.

0003895430 IP ACCESSION NO: 10980141 Chemistry of a new investigational quadrivalent meningococcal conjugate vaccine that is immunogenic at all ages

Broeker, Michael; Dull, Peter M; Rappuoli, Rino; ****Costantino, Paolo****

Novartis Vaccines and Diagnostics GmbH & Co. KG, Marburg, Germany, [mailto:Michael.Broeker@Novartis.com]

Vaccine, v 27, n 41, p 5574-5580, September 2009 PUBLICATION DATE: 2009

PUBLISHER: Elsevier Science, The Boulevard Langford Lane Kidlington Oxford OX5 1GB UK, [mailto:usinfo-f@elsevier.com], [URL:http://www.elsevier.nl]

DOCUMENT TYPE: Journal Article RECORD TYPE: Abstract LANGUAGE: English SUMMARY LANGUAGE: English ISSN: 0264-410X ELECTRONIC ISSN: 1873-2518 DOI: 10.1016/j.vaccine.2009.07.036

FILE SEGMENT: Bacteriology Abstracts (Microbiology B); Immunology Abstracts

ABSTRACT:

Meningococcal disease is a serious medical condition that can prove fatal within hours in otherwise healthy individuals. Disease incidence is highest in infants, yet there is no broadly protective quadrivalent vaccine that covers this age group. A new investigational quadrivalent meningococcal glycoconjugate vaccine against meningococcal serogroups A, C, W-135, and Y (MenACWY-CRM, Novartis Vaccines, Siena, Italy), has been developed to meet this medical need. This article discusses the vaccine technology behind MenACWY-CRM, focusing on the heritage of CRM sub(197), the conjugation chemistry, the sizing of the oligosaccharides, and the advantages that these may confer on the vaccine. We highlight the differences between available vaccines and look at the clinical experience with vaccines against other diseases, demonstrating the importance of each component to the immunogenicity of conjugate vaccines. The specific technological approach, including conjugation of meningococcal oligosaccharides of defined length to the CRM sub(197) protein, has led to a vaccine that has the potential to provide broad meningococcal protection against serogroups A, C, W-135, and Y for all ages.

10/7/20 (Item 3 from file: 24)
DIALOG(R)File 24:CSA Life Sciences Abstracts
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0003382995 IP ACCESSION NO: 8554501

Physicochemical characterisation of glycoconjugate vaccines for prevention of meningococcal diseases

Bardotti, Angela, Averani, Giovanni; ****Berti, Francesco****; Berti, Stefania; Carinci, Valeria; D'Ascenzi, Sandro; Fabbri, Barbara; Giannini, Sara; Giannozzi, Aldo; Magagnoll, Claudia; Proietti, Daniela; Norelli, Francesco; Rappuoli, Rino; Ricci, Stefano; ****Costantino, Paolo****

Novartis Vaccines and Diagnostics Srl, Via Fiorentina 1, 53100 Siena, Italy, [mailto:paolo.costantino@novartis.com]

Vaccine, v 26, n 18, p 2284-2296, April 2008 PUBLICATION DATE: 2008

PUBLISHER: Elsevier Science, The Boulevard Langford Lane Kidlington Oxford

OX5 1GB UK, [mailto:usinfo-f@elsevier.com], [URL:http://www.elsevier.nl]

DOCUMENT TYPE: Journal Article RECORD TYPE: Abstract LANGUAGE: English SUMMARY LANGUAGE: English ISSN: 0264-410X ELECTRONIC ISSN: 1873-2518 DOI: 10.1016/i.vaccine.2008.01.022

FILE SEGMENT: Bacteriology Abstracts (Microbiology B); Immunology Abstracts

Bacterial ****capsular**** polysaccharides covalently linked to an appropriate carrier protein represent the best tool to induce a protective immune response against a wide range of bacterial diseases, such as meningococcal infections. We describe here the physico-chemical characterisation of glycoconjugate molecules designed to prepare a vaccine against ****Neisseria**** ****meningitidis**** serogroups A, C, W135 and Y. The use of a selective conjugation chemistry resulted in well characterised, reproducible and traceable glycoconjugate that can be consistently manufactured at large scale. A pool of physical and spectroscopic methods was used to establish glycosylation ratio, identity, molecular weight profiles, integrity of carrier protein and sites of glycosylation, assuring effective and consistent lots of vaccines.

10/7/21 (Item 4 from file: 24) DIALOG(R)File 24:CSA Life Sciences Abstracts (c) 2011 CSA. All rts. reserv.

0003149696 IP ACCESSION NO: 7935788 Introduction of Zwitterionic Motifs into Bacterial Polysaccharides Generates TLR2 Agonists Able to Activate APCs

Gallorini, Simona; ****Berti, Francesco****; Parente, Pierino; Baronio, Roberta; Aprea, Susanna; D'Oro, Ugo; Pizza, Mariagrazia; Telford, John L; Wack, Andreas

Novartis Vaccines Research Center, Siena, Italy

Journal of Immunology, v 179, n 12, p 8208-8215, December 2007 PUBLICATION DATE: 2007

PUBLISHER: American Association of Immunologists, 9650 Rockville Pike Bethesda MD 20814-3998 USA, [URL:http://www.jimmunol.org/]

DOCUMENT TYPE: Journal Article RECORD TYPE: Abstract LANGUAGE: English SUMMARY LANGUAGE: English ISSN: 0022-1767

FILE SEGMENT: Bacteriology Abstracts (Microbiology B); Immunology Abstracts

ABSTRACT:

It was shown previously that bacterial polysaccharides (PS), which naturally contain both positive and negative charges, are able to activate T cells and APCs. However, the vast majority of bacterial PS are anionic and do not have these properties. In this study, we show that chemical introduction of positive charges into naturally anionic bacterial PS confers to the resulting zwitterionic PS (ZPS) the ability to activate pure human monocytes, monocyte-derived dendritic cells, and mouse bone marrow-derived dendritic cells, as do natural bacterial ZPS. Cells are induced to up-regulate MHC class II and costimulatory molecules and to

produce cytokines. In mixed monocyte-T cell cocultures, ZPS induce MHC II-dependent T cell proliferation and up-regulation of activation markers. These stimulatory qualities of ZPS disappear when the positive charge is chemically removed from the molecules and thus the zwitterionic motif is destroyed. The ability of natural and chemically derived ZPS to activate

```
APCs can be blocked by anti-TLR2 mAbs, and TLR2 transfectants show reporter
gene transcription upon incubation with ZPS. In conclusion, the generation
of a zwitterionic motif in bacterial PS confers the ability to activate
both APCs and T cells. This finding has important implications for the
design of novel ****polysaccharide**** vaccines.
10/7/22
            (Item 1 from file: 399)
DIALOG(R)File 399:CA SEARCH(R)
(c) 2011 American Chemical Society. All rts. reserv.
 154158714
             CA: 154(8)158714u
                                  JOURNAL
 First Synthesis of C. difficile PS-II Cell Wall Polysaccharide Repeating
 Unit.
 AUTHOR(S): Danieli, Elisa; Lay, Luigi; Proietti, Daniela; Berti,
Francesco; Costantino, Paolo; Adamo, Roberto
 LOCATION: Vaccine Chemistry Department, Novartis Vaccines & Diagnostics,
53100, Siena, Italy
 JOURNAL: Organic Lett. (Organic Letters) DATE: 2011 VOLUME: 13 NUMBER: 3
 PAGES: 378-381 CODEN: ORLEF7 MEDIA TYPE: online computer file ISSN:
1523-7052 LANGUAGE: English PUBLISHER: American Chemical Society
 SECTION:
   CA233008 Carbohydrates
 IDENTIFIERS: Clostridium difficile surface polysaccharide repeating unit
   prepn
 DESCRIPTORS:
Clostridium difficile... Glycosides... Glycosylation... Oligosaccharides...
   convergent synthesis of C. difficile surface polysaccharide repeating
   unit and its nonphosphorylated analog
Diarrhea...
   nosocomial; convergent synthesis of C. difficile surface polysaccharide
   repeating unit and its nonphosphorylated analog
 CAS REGISTRY NUMBERS:
34637-22-4 108869-64-3 121238-27-5 278784-83-1 1010440-34-2
   1236190-06-9P 1256157-14-8P 1262208-34-3P 1262208-37-6P
   1262208-39-8P 1262208-42-3P 1262208-43-4P 1262208-45-6P
   1262208-49-0P 1262208-52-5P 1262208-53-6P 1262208-54-7P
   1262208-55-8P 1262208-57-0P 1262208-58-1P 1262208-59-2P
   1262208-61-6P 1262208-63-8P 1262208-65-0P 1262208-67-2P
   1262208-69-4P 1262208-71-8P 1262208-73-0P 1262208-75-2P convergent
   synthesis of C. difficile surface polysaccharide repeating unit and its
   nonphosphorvlated analog
            (Item 2 from file: 399)
10/7/23
DIALOG(R)File 399:CA SEARCH(R)
(c) 2011 American Chemical Society. All rts. reserv.
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153478604 CA: 153(19)478604w Combinations including pneumococcal serotype 14 saccharide INVENTOR (AUTHOR): Costantino, Paolo LOCATION: Switz. ASSIGNEE: Novartis A.-G. PATENT: PCT International ; WO 2010109325 A2 DATE: 20100930 APPLICATION: WO 2010IB735 (20100324) *US 2009PV162996 (20090324) PAGES: 47pp. CODEN: PIXXD2 LANGUAGE: English PATENT CLASSIFICATIONS:

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IPCR/8 + Level Value Position Status Version Action Source Office:
                       A I F B 20060101
      A61K-0039/02
  DESIGNATED COUNTRIES: AE; AG; AL; AM; AO; AT; AU; AZ; BA; BB; BG; BH; BR;
BW; BY; BZ; CA; CH; CL; CN; CO; CR; CU; CZ; DE; DK; DM; DO; DZ; EC; EE; EG;
ES; FI; GB; GD; GE; GH; GM; GT; HN; HR; HU; ID; IL; IN; IS; JP; KE; KG; KM;
KN; KP; KR; KZ; LA; LC; LK; LR; LS; LT; LU; LY; MA; MD; ME; MG; MK; MN; MW;
MX; MY; MZ; NA; NG; NI; NO; NZ; OM; PE; PG; PH; PL; PT; RO; RS; RU; SC; SD;
SE; SG; SK; SL; SM; ST; SV; SY; TH; TJ DESIGNATED REGIONAL: AT; BE; BG; CH
; CY; CZ; DE; DK; EE; ES; FI; FR; GB; GR; HR; HU; IE; IS; IT; LT; LU; LV;
MC; MK; MT; NL; NO; PL; PT; RO; SE; SI; SK; SM; TR; BF; BJ; CF; CG; CI; CM;
GA; GN; GQ; GW; ML; MR; NE; SN; TD; TG; BW; GH; GM; KE; LR; LS; MW; MZ; NA;
SD; SL; SZ; TZ; UG; ZM; ZW; AM; AZ; BY; KG; KZ; MD; RU; TJ; TM
  SECTION:
    CA215002 Immunochemistry
    CA214XXX Mammalian Pathological Biochemistry
  IDENTIFIERS: Streptococcus Neisseria combination vaccine antigen
    crossreactivity
  DESCRIPTORS:
Polysaccharides...
    capsular; combination pneumococcal meningococcal vaccine with reduced
    lacto-N-neotetraose antigenicity
Development, mammalian postnatal ...
    child; combination pneumococcal meningococcal vaccine with reduced
    lacto-N-neotetraose antigenicity
Streptococcus pneumoniae... Vaccines... Antibodies and Immunoglobulins...
Human... Mammalia...
    combination pneumococcal meningococcal vaccine with reduced
    lacto-N-neotetraose antigenicity
Proteins...
    complement factor H-binding; of combination pneumococcal meningococcal
    vaccine with reduced lacto-N-neotetraose antigenicity
Protein D...
    conjugates; with capsular polysaccharides of Streptococcus pneumoniae
Toxoids...
    diphtheria, conjugates; with capsular polysaccharides of Streptococcus
    pneumoniae
Toxins...
    diphtheria, fragment; conjugates with capsular polysaccharides of
    Streptococcus pneumoniae
Lipid A...
    for conjugation of carrier proteins to meningococcal
    lipooligosaccharides
Protein sequences...
    for factor H-binding proteins of Neisseria meningitidis
Neisseria meningitidis...
    group B; combination pneumococcal meningococcal vaccine with reduced
    lacto-N-neotetraose antigenicity
Lipopolysaccharides...
    lipooligosaccharides; combination pneumococcal meningococcal vaccine
    with reduced lacto-N-neotetraose antigenicity
Transport proteins...
    TbpA; meningococcal lipooligosaccharides derived from strains with
```

vesicles; combination pneumococcal meningococcal vaccine with reduced CAS REGISTRY NUMBERS: 7784-30-7 adjuvant for combination pneumococcal meningococcal vaccine with reduced lacto-N-neotetraose antigenicity

tetanus, conjugates; with capsular polysaccharides of Streptococcus

over-expression of

lacto-N-neotetraose antigenicity

Toxoids...

pneumoniae Cell membrane...

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1246898-01-0 1246898-02-1 1246898-03-2 amino acid sequence; of
   combination pneumococcal meningococcal vaccine with reduced
    lacto-N-neotetraose antigenicity
13007-32-4 combination pneumococcal meningococcal vaccine with reduced
    lacto-N-neotetraose antigenicity
10149-14-1 for conjugation of carrier proteins to meningococcal
    lipooligosaccharides
9033-07-2 Lacto-N-neotetraose biosynthesis glycosyl transferase LqtB;
   meningococcal lipooligosaccharides derived from strains deficient for
9032-89-7 37277-64-8 219610-16-9 meningococcal lippoligosaccharides
   derived from strains deficient for
7429-90-5D salts, adjuvant for combination pneumococcal meningococcal
   vaccine with reduced lacto-N-neotetraose antigenicity
1246900-96-8 1246900-97-9 1246900-98-0 1246900-99-1 1246901-00-7
   1246901-01-8 1246901-02-9 1246901-03-0 1246901-04-1 1246901-05-2
   1246901-06-3 1246901-07-4 1246901-08-5 1246901-09-6 1246901-10-9
   1246901-11-0 1246901-12-1 1246901-13-2 1246901-14-3 1246901-15-4
   1246901-16-5 1246901-17-6 1246901-18-7 1246901-19-8 1246901-20-1
   1246901-21-2 1246901-22-3 1246901-23-4 1246901-24-5 1246901-25-6
   1246901-26-7 1246901-27-8 1246901-28-9 1246901-29-0 1246901-30-3
   1246901-31-4 1246901-32-5 1246901-33-6 1246901-34-7 1246901-35-8
    1246901-36-9 1246901-37-0 1246901-38-1 1246901-39-2 1246901-40-5
    1246901-41-6 1246901-42-7 1246901-43-8 1246901-44-9 1246901-45-0
   1246901-47-2 unclaimed protein sequence; combinations including
   pneumococcal serotype 14 saccharide
219724-66-0 1112044-56-0 64134-30-1 1246901-46-1 848652-29-9 unclaimed
   sequence; combinations including pneumococcal serotype 14 saccharide
10/7/24
            (Item 3 from file: 399)
DIALOG(R)File 399:CA SEARCH(R)
(c) 2011 American Chemical Society. All rts. reserv.
             CA: 152(24)543993t
 152543993
                                   PATENT
 Purification of Streptococcus pyogenes GAS carbohydrate by anionic
 exchange chromatography
 INVENTOR (AUTHOR): Costantino, Paolo; Berti, Francesco; Kabanova, Anna;
Romano, Maria Rosaria
 LOCATION: Switz.
 ASSIGNEE: Novartis AG
 PATENT: PCT International ; WO 201049806 A1 DATE: 20100506
 APPLICATION: WO 20091B7346 (20091027) *US 2008PV108763 (20081027)
 PAGES: 56pp. CODEN: PIXXD2 LANGUAGE: English
 PATENT CLASSIFICATIONS:
   IPCR/8 + Level Value Position Status Version Action Source Office:
     B01D-0015/36 A I F B 20060101 H EP
     B01D-0061/14
                      A I L B 20060101
                                                      H EP
     C07K-0001/34
                      A I L B 20060101
                                                     H EP
                       A I L B 20060101
                                                     H EP
     A61K-0039/09
     C07K-0001/18
                      A I L B 20060101
A I L B 20060101
                                                      H EP
                                                      H EP
     C12P-0019/00
 DESIGNATED COUNTRIES: AE; AG; AL; AM; AO; AT; AU; AZ; BA; BB; BG; BH; BR;
BW; BY; BZ; CA; CH; CL; CN; CO; CR; CU; CZ; DE; DK; DM; DO; DZ; EC; EE; EG;
ES; FI; GB; GD; GE; GH; GM; GT; HN; HR; HU; ID; IL; IN; IS; JP; KE; KG; KM;
KN; KP; KR; KZ; LA; LC; LK; LR; LS; LT; LU; LY; MA; MD; ME; MG; MK; MN; MW;
MX; MY; MZ; NA; NG; NI; NO; NZ; OM; PE; PG; PH; PL; PT; RO; RS; RU; SC; SD;
SE; SG; SK; SL; SM; ST; SV; SY; TJ; TM DESIGNATED REGIONAL: AT; BE; BG; CH
; CY; CZ; DE; DK; EE; ES; FI; FR; GB; GR; HR; HU; IE; IS; IT; LT; LU; LV;
MC; MK; MT; NL; NO; PL; PT; RO; SE; SI; SK; SM; TR; BF; BJ; CF; CG; CI; CM;
GA; GN; GQ; GW; ML; MR; NE; SN; TD; TG; BW; GH; GM; KE; LS; MW; MZ; NA; SD;
SL; SZ; TZ; UG; ZM; ZW; AM; AZ; BY; KG; KZ; MD; RU; TJ; TM
 SECTION:
```

CA209003 Biochemical Methods

IDENTIFIERS: Streptococcus GAS cell wall polysaccharide purifn, anionic exchange chromatog Streptococcus GAS cell wall polysaccharide DESCRIPTORS:

Polysaccharides...

GASP (group A Streptococcus cell wall polysaccharide); purification of Streptococcus pyogenes GAS carbohydrate by anionic exchange chromatog.

mobile phase buffer; purification of Streptococcus pyogenes GAS carbohydrate by anionic exchange chromatog.

Filtration...

orthogonal; purification of Streptococcus pyogenes GAS carbohydrate by anionic exchange chromatog.

Anion exchange chromatography... Filtration... Nucleic acids... Proteins... Size-exclusion chromatography... Streptococcus group A... Streptococcus pyogenes... Ultrafiltration...

purification of Streptococcus pyogenes GAS carbohydrate by anionic exchange chromatog. Filtration...

tangential-flow filtration; purification of Streptococcus pyogenes GAS carbohydrate by anionic exchange chromatog. CAS REGISTRY NUMBERS:

334756-67-1 anion exchange matrix; purification of Streptococcus pyogenes GAS

carbohydrate by anionic exchange chromatog.

9004-61-9 118214-04-3 purification of Streptococcus pyogenes GAS carbohydrate by anionic exchange chromatog. 9004-54-0 uses, gel filtration on; purification of Streptococcus pyogenes GAS

carbohydrate by anionic exchange chromatog.

64-17-5 uses, mobile phase buffer; purification of Streptococcus pyogenes GAS carbohydrate by anionic exchange chromatog.

10/7/25 (Item 4 from file: 399) DIALOG(R)File 399:CA SEARCH(R) (c) 2011 American Chemical Society, All rts. reserv.

152523448 CA: 152(23)523448r JOURNAL

Neisseria meningitidis GNA2132, a heparin-binding protein that induces protective immunity in humans

AUTHOR(S): Serruto, Davide; Spadafina, Tiziana; Ciucchi, Laura; Lewis, Lisa A.; Ram, Sanjav; Tontini, Marta; Santini, Laura; Biolchi, Alessia; Seib, Kate L.; Giuliani, Marzia M.; Donnelly, John J.; Berti, Francesco; Savino, Silvana; Scarselli, Maria; Costantino, Paolo; Kroll, J. Simon; O'Dwyer, Cliona; Qui, Jiazhou; Plaut, Andrew G.; Moxon, Richard; Rappuoli, Rino; Pizza, Mariagrazia; Arico, Beatrice

LOCATION: Novartis Vaccines and Diagnostics, 53100, Siena, Italy

JOURNAL: Proc. Natl. Acad. Sci. U. S. A. (Proceedings of the National Academy of Sciences of the United States of America) DATE: 2010 VOLUME: 107 NUMBER: 8 PAGES: 3770-3775, S3770/1-S3770/9 CODEN: PNASA6 ISSN: 0027-8424 LANGUAGE: English PUBLISHER: National Academy of Sciences

CA215002 Immunochemistry

CA210XXX Microbial, Algal, and Fungal Biochemistry IDENTIFIERS: Neisseria GNA2132 protein immunity

DESCRIPTORS:

Protein motifs...

arginine-rich; in heparin-binding activity of GNA2132 of Neisseria Lactoferrins...

cleavage of neisserial heparin-binding antigen by

Proteoglycans...

heparitin sulfate-containing; GNA2132 protein of Neisseria meningitidis binds to

```
Transport proteins...
    NalP; cleavage of neisserial heparin-binding antigen by
Human... Neisseria meningitidis...
    Neisseria meningitidis GNA2132 is heparin-binding protein that induces
    protective immunity in humans
Virulence (microbial) ...
    neisserial heparin-binding antigen as factor in
    NHBA (neisserial heparin-binding antigen); Neisseria meningitidis
    GNA2132 is heparin-binding protein that induces protective immunity in
    humans
  CAS REGISTRY NUMBERS:
9005-49-6 biological studies, GNA2132 protein of Neisseria meningitidis
    binds to
 10/7/26
             (Item 5 from file: 399)
DIALOG(R)File 399:CA SEARCH(R)
(c) 2011 American Chemical Society. All rts. reserv.
  152376270
             CA: 152(17)376270w
  Analysis of Vi saccharide of Salmonella typhi
  INVENTOR(AUTHOR): Berti, Francesco; Micoli, Francesca; Proietti, Daniela
  LOCATION: Italy
  ASSIGNEE: Novartis Vaccines and Diagnostics S.R.L.
  PATENT: Ital. Appl.; IT 2008MI001079 A1 DATE: 20080913
  APPLICATION: IT 2008MI1079 (20080613)
  PAGES: 36pp. CODEN: ITXXCZ LANGUAGE: English
  PATENT CLASSIFICATIONS:
   CLASS: G01N-000/A
  SECTION:
   CA209003 Biochemical Methods
  IDENTIFIERS: Vi saccharide Salmonella NMR anion exchange HPLC
  DESCRIPTORS:
Electrochemical sensors...
    amperometric, pulsed; in determination of Vi saccharide of Salmonella typhi by
    anion exchange HPLC
Polysaccharides...
    capsular, of Salmonella typhi, deacetylated; determination of Vi saccharide of
    Salmonella typhi by NMR and anion exchange HPLC
Citrobacter freundii...
    determination of Vi saccharide by NMR and anion exchange HPLC
Salmonella typhi... NMR spectroscopy... Anion exchange HPLC...
    determination of Vi saccharide of Salmonella typhi by NMR and anion exchange
    HPLC
Antigens...
    Vi; determination of Vi saccharide of Salmonella typhi by NMR and anion
    exchange HPLC
  CAS REGISTRY NUMBERS:
1310-73-2 analysis, deacetylation of Vi saccharide of Salmonella typhi by
76-05-1 analysis, hydrolysis of Vi saccharide of Salmonella typhi by
14014-06-3 deacetylation of Vi saccharide of Salmonella typhi by
77-92-9 64-17-5 uses, internal reference standard for determination of Vi
saccharide of
    Salmonella typhi by NMR
 10/7/27
             (Item 6 from file: 399)
DIALOG(R)File 399:CA SEARCH(R)
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```

152045529 CA: 152(2)45529c PATENT

```
Conjugated Vi saccharides for vaccines against typhoid fever
  INVENTOR (AUTHOR): Micoli, Francesca; Costantino, Paolo; Berti, Francesco
  LOCATION: Switz.
  ASSIGNEE: Novartis AG
  PATENT: PCT International ; WO 2009150543 A2 DATE: 20091217
  APPLICATION: WO 2009IB6285 (20090612) *GB 200810894 (20080613)
  PAGES: 42pp. CODEN: PIXXD2 LANGUAGE: English
  PATENT CLASSIFICATIONS:
    IPCR/8 + Level Value Position Status Version Action Source Office:
      A61K-0047/48
                       A I F B 20060101
      A61P-0043/00
                       A N L B 20060101
                                                        H EP
  DESIGNATED COUNTRIES: AE; AG; AL; AM; AO; AT; AU; AZ; BA; BB; BG; BH; BR;
BW; BY; BZ; CA; CH; CL; CN; CO; CR; CU; CZ; DE; DK; DM; DO; DZ; EC; EE; EG;
ES; FI; GB; GD; GE; GH; GM; GT; HN; HR; HU; ID; IL; IN; IS; JP; KE; KG; KM;
KN; KP; KR; KZ; LA; LC; LK; LR; LS; LT; LU; LY; MA; MD; ME; MG; MK; MN; MW;
MX; MY; MZ; NA; NG; NI; NO; NZ; OM; PE; PG; PH; PL; PT; RO; RS; RU; SC; SD;
SE; SG; SK; SL; SM; ST; SV; SY; TJ; TM DESIGNATED REGIONAL: AT; BE; BG; CH
; CY; CZ; DE; DK; EE; ES; FI; FR; GB; GR; HR; HU; IE; IS; IT; LT; LU; LV;
MC; MK; MT; NL; NO; PL; PT; RO; SE; SI; SK; TR; BF; BJ; CF; CG; CI; CM; GA;
GN; GQ; GW; ML; MR; NE; SN; TD; TG; BW; GH; GM; KE; LS; MW; MZ; NA; SD; SL;
SZ; TZ; UG; ZM; ZW; AM; AZ; BY; KG; KZ; MD; RU; TJ; TM
  SECTION:
   CA263003 Pharmaceuticals
    CA215XXX Immunochemistry
  IDENTIFIERS: Salmonella capsular polysaccharide Vi protein conjugate
   typhoid vaccine
  DESCRIPTORS:
Polysaccharides...
    capsular, Vi; conjugates of Vi Salmonella polysaccharides with carrier
    proteins for typhoid vaccines
Salmonella typhi... Typhoid fever... Antigens... Immune adjuvants...
Pharmaceutical carriers... Drug delivery systems...
    conjugates of Vi Salmonella polysaccharides with carrier proteins for
    typhoid vaccines
Polysaccharides...
```

conjugates, with carrier proteins; conjugates of Vi Salmonella polysaccharides with carrier proteins for typhoid vaccines

Toxoids... tetanus, conjugates with Vi polysaccharide; conjugates of Vi Salmonella polysaccharides with carrier proteins for typhoid vaccines

typhoid fever; conjugates of Vi Salmonella polysaccharides with carrier proteins for typhoid vaccines

CAS REGISTRY NUMBERS:

DIALOG(R)File 399:CA SEARCH(R)

LOCATION: Switz.

10/7/28

1892-57-5 conjugates of Vi Salmonella polysaccharides with carrier proteins for typhoid vaccines

600173-37-3DP conjugates with Vi Salmonella polysaccharides, conjugates of Vi Salmonella polysaccharides with carrier proteins for typhoid vaccines

1071-93-8 linker; conjugates of Vi Salmonella polysaccharides with carrier proteins for typhoid vaccines

```
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151156623 CA: 151(7)156623c PATENT
Lipopolysaccharide decontamination during the purification of biopharmaceuticals
INVENTOR (AUTHOR): Costantino, Paolo
```

(Item 7 from file: 399)

```
ASSIGNEE: Novartis AG
 PATENT: PCT International ; WO 200987571 A2 DATE: 20090716
 APPLICATION: WO 2009IB133 (20090107) *GB 2008228 (20080107)
 PAGES: 13pp. CODEN: PIXXD2 LANGUAGE: English
 PATENT CLASSIFICATIONS:
   IPCR/8 + Level Value Position Status Version Action Source Office:
     B01J-0020/26 A I F B 20060101
                                                    H EP
     B01J-0020/28
                      A I L B 20060101
                                                      H EP
     B01D-0061/00
                      A I L B 20060101
                                                      H EP
     B01D-0067/00
                      A I L B 20060101
                                                      H EP
 DESIGNATED COUNTRIES: AE; AG; AL; AM; AO; AT; AU; AZ; BA; BB; BG; BH; BR;
BW; BY; BZ; CA; CH; CN; CO; CR; CU; CZ; DE; DK; DM; DO; DZ; EC; EE; EG; ES;
FI; GB; GD; GE; GH; GM; GT; HN; HR; HU; ID; IL; IN; IS; JP; KE; KG; KM; KN;
KP; KR; KZ; LA; LC; LK; LR; LS; LT; LU; LY; MA; MD; ME; MG; MK; MN; MW; MX;
MY; MZ; NA; NG; NI; NO; NZ; OM; PG; PH; PL; PT; RO; RS; RU; SC; SD; SE; SG;
SK; SL; SM; ST; SV; SY; TJ; TM; TN; TR DESIGNATED REGIONAL: AT; BE; BG; CH
; CY; CZ; DE; DK; EE; ES; FI; FR; GB; GR; HR; HU; IE; IS; IT; LT; LU; LV;
MC; MK; MT; NL; NO; PL; PT; RO; SE; SI; SK; TR; BF; BJ; CF; CG; CI; CM; GA;
GN; GQ; GW; ML; MR; NE; SN; TD; TG; BW; GH; GM; KE; LS; MW; MZ; NA; SD; SL;
SZ; TZ; UG; ZM; ZW; AM; AZ; BY; KG; KZ; MD; RU; TJ; TM
 SECTION:
   CA263008 Pharmaceuticals
   CA210XXX Microbial, Algal, and Fungal Biochemistry
 IDENTIFIERS: lipopolysaccharide decontamination biopharmaceutical purifn
 DESCRIPTORS:
Drugs...
   biopharmaceuticals; lipopolysaccharide decontamination during the
   purification of biopharmaceuticals
Purification... Decontamination... Lipopolysaccharides... Polymers...
Gram-negative bacteria... Proteobacteria... Cyanobacteria... Spirochaeta...
Green sulfur bacteria... Chloroflexi... Crenarchaeota... Eubacteria...
Bacilli... Membranes, nonbiological...
   lipopolysaccharide decontamination during the purification of
   biopharmaceuticals
 CAS REGISTRY NUMBERS:
25067-34-9 lipopolysaccharide decontamination during purification of
   biopharmaceuticals
71927-65-6 1069-03-0 lipopolysaccharide decontamination during the
   purification of biopharmaceuticals
10/7/29
            (Item 8 from file: 399)
DIALOG(R)File 399:CA SEARCH(R)
(c) 2011 American Chemical Society. All rts. reserv.
 151099686
              CA: 151(5)99686z
                                  PATENT
 Production and purification of Streptococcus agalactiae capsular
 polysaccharides
 INVENTOR(AUTHOR): Costantino, Paolo; Norelli, Francesco; Berti, Francesco
; Cicala, Concetta Maria; Bazzocchi, Giulia; Fontani, Silvia; Olivieri,
Roberto
 LOCATION: Switz.
 ASSIGNEE: Novartis A .- G.
 PATENT: PCT International ; WO 200981276 A2 DATE: 20090702
 APPLICATION: WO 20081B3729 (20081219) *US 2007PV8941 (20071220) *GB
200818453 (20081008)
 PAGES: 159pp. CODEN: PIXXD2 LANGUAGE: English
 PATENT CLASSIFICATIONS:
   IPCR/8 + Level Value Position Status Version Action Source Office:
     C12N-0001/20 A I F B 20060101 H EP
     A61K-0039/04
                                                     H EP
                     A I L B 20060101
                                                    H EP
     C12P-0019/04
                     A I L B 20060101
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DESIGNATED COUNTRIES: AE; AG; AL; AM; AO; AT; AU; AZ; BA; BB; BG; BH; BR;
BW; BY; BZ; CA; CH; CN; CO; CR; CU; CZ; DE; DK; DM; DO; DZ; EC; EE; EG; ES;
FI; GB; GD; GE; GH; GM; GT; HN; HR; HU; ID; IL; IN; IS; JP; KE; KG; KM; KN;
KP; KR; KZ; LA; LC; LK; LR; LS; LT; LU; LY; MA; MD; ME; MG; MK; MN; MW; MX;
MY; MZ; NA; NG; NI; NO; NZ; OM; PG; PH; PL; PT; RO; RS; RU; SC; SD; SE; SG;
SK; SL; SM; ST; SV; SY; TJ; TM; TN; TR DESIGNATED REGIONAL: AT; BE; BG; CH
; CY; CZ; DE; DK; EE; ES; FI; FR; GB; GR; HR; HU; IE; IS; IT; LT; LU; LV;
MC; MT; NL; NO; PL; PT; RO; SE; SI; SK; TR; BF; BJ; CF; CG; CI; CM; GA; GN;
GO; GW; ML; MR; NE; SN; TD; TG; BW; GH; GM; KE; LS; MW; MZ; NA; SD; SL; SZ;
TZ; UG; ZM; ZW; AM; AZ; BY; KG; KZ; MD; RU; TJ; TM
  SECTION:
    CA216002 Fermentation and Bioindustrial Chemistry
    CA210XXX Microbial, Algal, and Fungal Biochemistry
  IDENTIFIERS: Streptococcus capsular polysaccharide fermn purifn
  DESCRIPTORS:
Lipopolysaccharides...
    bacterial; production and purification of Streptococcus agalactiae capsular
    polysaccharides
Polysaccharides...
    capsular; production and purification of Streptococcus agalactiae capsular
    polysaccharides
Detergents...
    cationic; production and purification of Streptococcus agalactiae capsular
```

polysaccharides Culture media...

defined; production and purification of Streptococcus agalactiae capsular polysaccharides

Yeast...

extract; production and purification of Streptococcus agalactiae capsular polysaccharides

Fermentation...

fed-batch; production and purification of Streptococcus agalactiae capsular polysaccharides Growth, microbial ...

kinetics; production and purification of Streptococcus agalactiae capsular

polysaccharides Filtration...

microfiltration; production and purification of Streptococcus agalactiae capsular polysaccharides

Acetylation... N-; production and purification of Streptococcus agalactiae capsular

polysaccharides Streptococcus agalactiae... pH... Temperature effects, biological... Mineral

elements... Vitamins... Amino acids... Filtration... Precipitation(chemical)... Solubilization... Ultrafiltration... Nucleic

acids... Vaccines... Culture media... Proteins... Pressure...

Nutrition, microbial... Centrifugation... Dialysis... Sterilization and Disinfection... Sialic acids...

production and purification of Streptococcus agalactiae capsular polysaccharides Chemical engineering design ...

scale-up; production and purification of Streptococcus agalactiae capsular polysaccharides

CAS REGISTRY NUMBERS:

7782-44-7 processes, dissolved; production and purification of Streptococcus agalactiae capsular polysaccharides

50-99-7 83-88-5 59-30-3 7647-14-5 56-41-7 74-79-3 56-85-9 56-40-6 71-00-1 73-32-5 61-90-5 56-87-1 63-68-3 63-91-2 147-85-3 56-45-1 72-19-5 73-22-3 72-18-4 56-84-8 56-86-0 60-18-4 processes, production

and purification of Streptococcus agalactiae capsular polysaccharides 58-85-5 98-92-0 137-08-6 67-03-8 58-56-0 7778-77-0 118830-14-1

10049-21-5 7758-11-4 52-89-1 108-24-7 production and purification of Streptococcus agalactiae capsular polysaccharides

- 7440-44-0 uses, activated; production and purification of Streptococcus agalactiae capsular polysaccharides
- 64-17-5 10043-52-4 uses, production and purification of Streptococcus agalactiae capsular polysaccharides

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10/7/30
             (Item 9 from file: 399)
DIALOG(R)File 399:CA SEARCH(R)
(c) 2011 American Chemical Society. All rts. reserv.
  150166205 CA: 150(9)166205m
                                   PATENT
 Vaccines comprising saccharide antigen conjugated with
  hydroxyapatite-binding carrier protein to facilitate purification
  INVENTOR (AUTHOR): Bigio, Massimo; Averani, Giovanni; Norelli, Francesco;
Berti, Francesco; Bellucci, Cinzia
  LOCATION: Switz.
  ASSIGNEE: Novartis AG
  PATENT: PCT International; WO 200910877 A2 DATE: 20090122
  APPLICATION: WO 2008IB2690 (20080717) *GB 200713880 (20070717)
  PAGES: 54pp. CODEN: PIXXD2 LANGUAGE: English
  PATENT CLASSIFICATIONS:
    CLASS: A61K-000/A
  DESIGNATED COUNTRIES: AE; AG; AL; AM; AO; AT; AU; AZ; BA; BB; BG; BH; BR;
BW; BY; BZ; CA; CH; CN; CO; CR; CU; CZ; DE; DK; DM; DO; DZ; EC; EE; EG; ES;
FI; GB; GD; GE; GH; GM; GT; HN; HR; HU; ID; IL; IN; IS; JP; KE; KG; KM; KN;
KP; KR; KZ; LA; LC; LK; LR; LS; LT; LU; LY; MA; MD; ME; MG; MK; MN; MW; MX;
MY; MZ; NA; NG; NI; NO; NZ; OM; PG; PH; PL; PT; RO; RS; RU; SC; SD; SE; SG;
SK; SL; SM; ST; SV; SY; TJ; TM; TN; TR DESIGNATED REGIONAL: AT; BE; BG; CH
; CY; CZ; DE; DK; EE; ES; FI; FR; GB; GR; HR; HU; IE; IS; IT; LT; LU; LV;
MC; MT; NL; NO; PL; PT; RO; SE; SI; SK; TR; BF; BJ; CF; CG; CI; CM; GA; GN;
GQ; GW; ML; MR; NE; SN; TD; TG; BW; GH; GM; KE; LS; MW; MZ; NA; SD; SL; SZ;
TZ; UG; ZM; ZW; AM; AZ; BY; KG; KZ; MD; RU; TJ; TM
 SECTION:
    CA215002 Immunochemistry
    CA209XXX Biochemical Methods
    CA263XXX Pharmaceuticals
  IDENTIFIERS: saccharide antigen carrier protein conjugate hydroxyapatite
    binding vaccine purifn
  DESCRIPTORS:
Drug delivery systems... Drugs... Immunostimulants...
    adjuvants; vaccines comprising saccharide antigen conjugated with
    hydroxyapatite-binding carrier protein to facilitate purification
Purification...
    affinity; vaccines comprising saccharide antigen conjugated with
    hydroxyapatite-binding carrier protein to facilitate purification
Toxins...
    B; vaccines comprising saccharide antigen conjugated with
    hydroxyapatite-binding carrier protein to facilitate purification
Polysaccharides...
    capsular, desialylated; vaccines comprising saccharide antigen
    conjugated with hydroxyapatite-binding carrier protein to facilitate
```

purification Proteins...

carriers; vaccines comprising saccharide antigen conjugated with hydroxyapatite-binding carrier protein to facilitate purification Antigens... Proteins... Polysaccharides...

conjugates; vaccines comprising saccharide antigen conjugated with hydroxyapatite-binding carrier protein to facilitate purification Proteins...

contaminant; vaccines comprising saccharide antigen conjugated with hydroxyapatite-binding carrier protein to facilitate purification Pharmaceutical excipients... diluents; vaccines comprising saccharide antigen conjugated with hydroxyapatite-binding carrier protein to facilitate purification Toxoids...

diphtheria; vaccines comprising saccharide antigen conjugated with hydroxyapatite-binding carrier protein to facilitate purification

hydroxylapatite; vaccines comprising saccharide antigen conjugated with hydroxyapatite-binding carrier protein to facilitate purification Vaccines...

influenza; vaccines comprising saccharide antigen conjugated with hydroxyapatite-binding carrier protein to facilitate purification Biological transport...

iron; vaccines comprising saccharide antigen conjugated with hydroxyapatite-binding carrier protein to facilitate purification Toxins...

pertussis, protein; vaccines comprising saccharide antigen conjugated with hydroxyapatite-binding carrier protein to facilitate purification Buffers...

phosphate, vaccines comprising saccharide antigen conjugated with hydroxyapatite-binding carrier protein to facilitate purification Hemolysins...

pneumolysins; vaccines comprising saccharide antigen conjugated with hydroxyapatite-binding carrier protein to facilitate purification Epitopes...

poly-; N19; vaccines comprising saccharide antigen conjugated with hydroxyapatite-binding carrier protein to facilitate purification Epitopes...

poly-; vaccines comprising saccharide antigen conjugated with hydroxyapatite-binding carrier protein to facilitate purification Proteins...

PspA (pneumococcal surface protein A); vaccines comprising saccharide antigen conjugated with hydroxyapatite-binding carrier protein to facilitate purification

Proteins...

recombinant; vaccines comprising saccharide antigen conjugated with hydroxyapatite-binding carrier protein to facilitate purification Streptococcus agalactiae...

serotype V; vaccines comprising saccharide antigen conjugated with hydroxyapatite-binding carrier protein to facilitate purification Vaccines...

synthetic; vaccines comprising saccharide antigen conjugated with hydroxyapatite-binding carrier protein to facilitate purification Toxoids...

tetanus; vaccines comprising saccharide antigen conjugated with hydroxyapatite-binding carrier protein to facilitate purification Toxins...

toxin A; vaccines comprising saccharide antigen conjugated with hydroxyapatite-binding carrier protein to facilitate purification Vaccines.. Carbohydrates.. Glycoconjugates.. Antigens.. Carriers.. Affinity chromatography.. Neisseria meningitidis.. Outer membrane proteins.. Heat-shock proteins.. Cytokines.. Hormones, animal.. Lymphokines... Growth factors, animal.. Protein D... Haemophilus influenzae.. Clostridium difficile... Streptococcus pneumoniae.. Pseudomonas aeruginosa... Staphylococcus aureus... Enterococcus faecalis.. Enterococcus faecalis... Versinia enterocolitica... Vibrio cholerae.. Salmonella typhi... Linking agents... Pharmaceutical gels... Drug delivery systems... Pharmaceutical carriers... Infection... Bacterial infection... vaccines comprising saccharide antigen conjugated with

hydroxyapatite-binding carrier protein to facilitate purification pH...

6.5-7.5; vaccines comprising saccharide antigen conjugated with hydroxyapatite-binding carrier protein to facilitate purification

CAS REGISTRY NUMBERS:

7439-89-6P biological studies, transport; vaccines comprising saccharide antigen conjugated with hydroxyapatite-binding carrier protein to facilitate purification

7664-38-2D salts, buffer systems; vaccines comprising saccharide antigen conjugated with hydroxyapatite-binding carrier protein to facilitate purification

1306-06-5 600173-37-3P 9001-63-2 vaccines comprising saccharide antigen conjugated with hydroxyapatite-binding carrier protein to facilitate purification

10/7/31 (Item 10 from file: 399) DIALOG(R)File 399:CA SEARCH(R) (c) 2011 American Chemical Society. All rts. reserv. 150041363 CA: 150(3)41363e PATENT Formulation of meningitis vaccines INVENTOR (AUTHOR): Contorni, Mario; Costantino, Paolo LOCATION: Switz. ASSIGNEE: Novartis AG PATENT: PCT International ; WO 2008149238 A2 DATE: 20081211 APPLICATION: WO 2008IB2121 (20080604) *US 2007PV933235 (20070604) PAGES: 28pp. CODEN: PIXXD2 LANGUAGE: English PATENT CLASSIFICATIONS: CLASS: A61K-000/A DESIGNATED COUNTRIES: AE; AG; AL; AM; AO; AT; AU; AZ; BA; BB; BG; BH; BR; BW; BY; BZ; CA; CH; CN; CO; CR; CU; CZ; DE; DK; DM; DO; DZ; EC; EE; EG; ES; FI; GB; GD; GE; GH; GM; GT; HN; HR; HU; ID; IL; IN; IS; JP; KE; KG; KM; KN; KP; KR; KZ; LA; LC; LK; LR; LS; LT; LU; LY; MA; MD; ME; MG; MK; MN; MW; MX; MY; MZ; NA; NG; NI; NO; NZ; OM; PG; PH; PL; PT; RO; RS; RU; SC; SD; SE; SG; SK; SL; SM; SV; SY; TJ; TM; TN; TR; TT DESIGNATED REGIONAL: AT; BE; BG; CH ; CY; CZ; DE; DK; EE; ES; FI; FR; GB; GR; HR; HU; IE; IS; IT; LT; LU; LV; MC; MT; NL; NO; PL; PT; RO; SE; SI; SK; TR; BF; BJ; CF; CG; CI; CM; GA; GN; GQ; GW; ML; MR; NE; SN; TD; TG; BW; GH; GM; KE; LS; MW; MZ; NA; SD; SL; SZ; TZ; UG; ZM; ZW; AM; AZ; BY; KG; KZ; MD; RU; TJ; TM SECTION: CA263006 Pharmaceuticals CA215XXX Immunochemistry IDENTIFIERS: meningitis vaccine formulation DESCRIPTORS:

Immunostimulants...

adjuvants; formulation of meningitis vaccines

Human poliovirus...

antigens of, conjugates with saccharides; formulation of meningitis

vaccines

Pertussis...

conjugates with saccharides; formulation of meningitis vaccines Toxoids...

diphtheria, conjugates with saccharides; formulation of meningitis vaccines

Vaccines...

diphtheria-tetanus-acellular pertussis-inactivated polio virus; formulation of meningitis vaccines

Vaccines...

diphtheria-tetanus-acellular pertussis-inactivated polio

virus-Haemophilus influenzae type b; formulation of meningitis vaccines Human... Meningitis... Pharmaceutical emulsions... Stabilizing agents... Vaccines...

formulation of meningitis vaccines

Neisseria meningitidis...

group A, saccharides of, conjugates of; formulation of meningitis

vaccines Neisseria meningitidis... group C, saccharides of, conjugates of; formulation of meningitis Neisseria meningitidis... group W-135, saccharides of, conjugates of; formulation of meningitis vaccines Neisseria meningitidis... group Y, saccharides of, conjugates of; formulation of meningitis vaccines Hepatitis B antigens ... HBsAq (hepatitis B surface antigen), conjugates with saccharides; formulation of meningitis vaccines Neisseria meningitidis... saccharides of, conjugates of; formulation of meningitis vaccines Toxoids... tetanus, conjugates with saccharides; formulation of meningitis vaccines Antibodies and Immunoglobulins... to PRP; formulation of meningitis vaccines Haemophilus influenzae... type b, saccharides of, conjugates of; formulation of meningitis vaccines CAS REGISTRY NUMBERS: 600173-37-3D conjugates with saccharides, formulation of meningitis vaccines 7784-30-7 formulation of meningitis vaccines 10/7/32 (Item 11 from file: 399) DIALOG(R) File 399:CA SEARCH(R) (c) 2011 American Chemical Society. All rts. reserv. 149174110 CA: 149(8)174110w PATENT Preparation of modified protein-conjugated capsular oligosaccharides and polysaccharides in study of vaccine for Neisseria meningitidis INVENTOR(AUTHOR): Bardotti, Angela; Berti, Francesco; Costantino, Paolo LOCATION: Switz. ASSIGNEE: Novartis AG PATENT: PCT International ; WO 200884411 A2 DATE: 20080717 APPLICATION: WO 2008IB1116 (20080111) *GB 2007562 (20070111) PAGES: 84pp. CODEN: PIXXD2 LANGUAGE: English PATENT CLASSIFICATIONS: CLASS: C07H-000/A DESIGNATED COUNTRIES: AE; AG; AL; AM; AO; AT; AU; AZ; BA; BB; BG; BH; BR; BW; BY; BZ; CA; CH; CN; CO; CR; CU; CZ; DE; DK; DM; DO; DZ; EC; EE; EG; ES; FI; GB; GD; GE; GH; GM; GT; HN; HR; HU; ID; IL; IN; IS; JP; KE; KG; KM; KN; KP; KR; KZ; LA; LC; LK; LR; LS; LT; LU; LY; MA; MD; ME; MG; MK; MN; MW; MX; MY; MZ; NA; NG; NI; NO; NZ; OM; PG; PH; PL; PT; RO; RS; RU; SC; SD; SE; SG; SK; SL; SM; SV; SY; TJ; TM; TN; TR; TT DESIGNATED REGIONAL: AT; BE; BG; CH ; CY; CZ; DE; DK; EE; ES; FI; FR; GB; GR; HR; HU; IE; IS; IT; LT; LU; LV; MC; MT; NL; NO; PL; PT; RO; SE; SI; SK; TR; BF; BJ; CF; CG; CI; CM; GA; GN; GO; GW; ML; MR; NE; SN; TD; TG; BW; GH; GM; KE; LS; MW; MZ; NA; SD; SL; SZ; TZ; UG; ZM; ZW; AM; AZ; BY; KG; KZ; MD; RU; TJ; TM SECTION: CA215002 Immunochemistry CA201XXX Pharmacology CA210XXX Microbial, Algal, and Fungal Biochemistry

CA263XXX Pharmaceuticals IDENTIFIERS: diphtheria toxin vaccine immunization Neisseria meningitidis capsular polysaccharide, vaccine immunization Neisseria meningitidis

CA233XXX Carbohydrates

```
capsular polysaccharide prepn human antibacterial, aminodeoxy
    oligosaccharide capsular polysaccharide protein conjugate adjuvant
    antigen prepn
  DESCRIPTORS:
Immunostimulants...
    adjuvants; preparation of modified protein-conjugated capsular
    oligosaccharides and polysaccharides in study of vaccine for Neisseria
    meningitidis
Polysaccharides, biological studies...
    capsular; preparation of modified protein-conjugated capsular
    oligosaccharides and polysaccharides in study of vaccine for Neisseria
    meningitidis
Antibodies and Immunoglobulins...
    IgG; preparation of modified protein-conjugated capsular oligosaccharides
    and polysaccharides in study of vaccine for Neisseria meningitidis
Neisseria meningitidis... Antibacterial agents... Bacterial infection...
Antigens... Human... Immunization... Oligosaccharides, biological studies...
Drugs... Toxins... Vaccines...
    preparation of modified protein-conjugated capsular oligosaccharides and
    polysaccharides in study of vaccine for Neisseria meningitidis
  CAS REGISTRY NUMBERS:
1039052-04-4D acetylated, derivs., repeating unit; preparation of modified
    protein-conjugated capsular oligosaccharides and polysaccharides in
    study of vaccine for Neisseria meningitidis
1039052-05-5D 1039052-06-6D CRM 197 derivative, preparation of modified
    protein-conjugated capsular oligosaccharides and polysaccharides in
    study of vaccine for Neisseria meningitidis
600173-37-3 1039052-05-5P 1039052-06-6P preparation of modified
    protein-conjugated capsular oligosaccharides and polysaccharides in
    study of vaccine for Neisseria meningitidis
 10/7/33
             (Item 12 from file: 399)
DIALOG(R) File 399:CA SEARCH(R)
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  147474589
              CA: 147(22)474589y
                                   PATENT
  Separation of conjugated and unconjugated components in vaccines
  INVENTOR (AUTHOR): Berti, Francesco; Galletti, Bruno; Parente, Pierino;
Costantino, Paolo
  LOCATION: Italy
  ASSIGNEE: Novartis Vaccines and Diagnostics Srl
  PATENT: PCT International ; WO 2007122512 A2 DATE: 20071101
  APPLICATION: WO 2007IB1855 (20070321) *GB 20065757 (20060322)
  PAGES: 33pp. CODEN: PIXXD2 LANGUAGE: English
  PATENT CLASSIFICATIONS:
    CLASS: A61K-000/A
 DESIGNATED COUNTRIES: AE; AG; AL; AM; AT; AU; AZ; BA; BB; BG; BH; BR; BW;
BY; BZ; CA; CH; CN; CO; CR; CU; CZ; DE; DK; DM; DZ; EC; EE; EG; ES; FI; GB;
GD; GE; GH; GM; GT; HN; HR; HU; ID; IL; IN; IS; JP; KE; KG; KM; KN; KP; KR;
KZ; LA; LC; LK; LR; LS; LT; LU; LY; MA; MD; MG; MK; MN; MW; MX; MY; MZ; NA;
NG; NI; NO; NZ; OM; PG; PH; PL; PT; RO; RS; RU; SC; SD; SE; SG; SK; SL; SM;
SV; SY; TJ; TM; TN; TR; TT; TZ; UA; UG DESIGNATED REGIONAL: AT; BE; BG; CH
; CY; CZ; DE; DK; EE; ES; FI; FR; GB; GR; HU; IE; IS; IT; LT; LU; LV; MC;
MT; NL; PL; PT; RO; SE; SI; SK; TR; BF; BJ; CF; CG; CI; CM; GA; GN; GQ; GW;
ML; MR; NE; SN; TD; TG; BW; GH; GM; KE; LS; MW; MZ; NA; SD; SL; SZ; TZ; UG;
ZM; ZW; AM; AZ; BY; KG; KZ; MD; RU; TJ; TM
  SECTION:
    CA263003 Pharmaceuticals
```

IDENTIFIERS: antigen saccharide conjugate basic salt pptn vaccine quality

control DESCRIPTORS:

```
Salts, uses...
    basic, lyotropic; separation of conjugated and unconjugated saccharides
    using basic reagents for quality control of vaccines
Streptococcus agalactiae...
    capsular saccharides; separation of conjugated and unconjugated saccharides
    using basic reagents for quality control of vaccines
Polysaccharides, biological studies...
    conjugates; separation of conjugated and unconjugated saccharides using
    basic reagents for quality control of vaccines
Toxins...
    diphtheria; separation of conjugated and unconjugated saccharides using
    basic reagents for quality control of vaccines
Carbohydrates, biological studies... Polysaccharides, biological studies...
Glycoconjugates... Antigens... Vaccines... Sialic acids...
Precipitation(chemical)... Basicity... Quality control...
    separation of conjugated and unconjugated saccharides using basic reagents
    for quality control of vaccines
Toxoids...
    tetanus; separation of conjugated and unconjugated saccharides using basic
    reagents for quality control of vaccines
  CAS REGISTRY NUMBERS:
7664-38-2D acidic alkali metal salts, separation of conjugated and unconjugated
    saccharides using basic reagents for quality control of vaccines
7664-93-9D 64-19-7D 77-92-9D 87-69-4D alkali metal salts, separation of
    conjugated and unconjugated saccharides using basic reagents for
    quality control of vaccines
600173-37-3 7758-11-4 separation of conjugated and unconjugated saccharides
    using basic reagents for quality control of vaccines
 10/7/34
             (Item 13 from file: 399)
DIALOG(R)File 399:CA SEARCH(R)
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  146294161
             CA: 146(15)294161v
                                    PATENT
  Zwitterionization to convert T-independent into T-dependent immunogenic
  bacterial capsular saccharides for use as vaccines
  INVENTOR(AUTHOR): Telford, John; Berti, Francesco; Wack, Andreas
  LOCATION: Italy
  ASSIGNEE: Novartis Vaccines and Diagnostics S.r.l.
  PATENT: PCT International; WO 200723386 A2 DATE: 20070301
 APPLICATION: WO 20061B2833 (20060824) *GB 200517353 (20050824) *GB
20067738 (20060419)
  PAGES: 56pp. CODEN: PIXXD2 LANGUAGE: English
  PATENT CLASSIFICATIONS:
    CLASS: C07H-000/A
  DESIGNATED COUNTRIES: AE; AG; AL; AM; AT; AU; AZ; BA; BB; BG; BR; BW; BY;
BZ; CA; CH; CN; CO; CR; CU; CZ; DE; DK; DM; DZ; EC; EE; EG; ES; FI; GB; GD;
GE; GH; GM; HN; HR; HU; ID; IL; IN; IS; JP; KE; KG; KM; KN; KP; KR; KZ; LA;
LC; LK; LR; LS; LT; LU; LV; LY; MA; MD; MG; MK; MN; MW; MX; MY; MZ; NA; NG;
NI; NO; NZ; OM; PG; PH; PL; PT; RO; RS; RU; SC; SD; SE; SG; SK; SL; SM; SV;
SY; TJ; TM; TN; TR; TT; TZ; UA; UG; US DESIGNATED REGIONAL: AT; BE; BG; CH
; CY; CZ; DE; DK; EE; ES; FI; FR; GB; GR; HU; IE; IS; IT; LT; LU; LV; MC;
NL; PL; PT; RO; SE; SI; SK; TR; BF; BJ; CF; CG; CI; CM; GA; GN; GQ; GW; ML;
MR; NE; SN; TD; TG; BW; GH; GM; KE; LS; MW; MZ; NA; SD; SL; SZ; TZ; UG; ZM;
ZW; AM; AZ; BY; KG; KZ; MD; RU; TJ; TM
  SECTION:
    CA215002 Immunochemistry
   CA263XXX Pharmaceuticals
  IDENTIFIERS: zwitterionization bacterial capsular saccharide antigen T
    cell activation vaccine
  DESCRIPTORS:
```

T cell(lymphocyte)...

activation; zwitterionization to convert T-independent into T-dependent immunogenic bacterial capsular saccharides for use as vaccines

Infection...

bacterial; zwitterionization to convert T-independent into T-dependent immunogenic bacterial capsular saccharides for use as vaccines

Polysaccharides, biological studies...

capsular; zwitterionization to convert T-independent into T-dependent immunogenic bacterial capsular saccharides for use as vaccines Hydrolysis...

galactose unit; zwitterionization to convert T-independent into T-dependent immunogenic bacterial capsular saccharides for use as vaccines

Neisseria meningitidis...

group A; zwitterionization to convert T-independent into T-dependent immunogenic bacterial capsular saccharides for use as vaccines

Neisseria meningitidis...

group B; zwitterionization to convert T-independent into T-dependent immunogenic bacterial capsular saccharides for use as vaccines Neisseria meningitidis...

ersseria meningitidis..

group C; zwitterionization to convert T-independent into T-dependent immunogenic bacterial capsular saccharides for use as vaccines Neisseria meningitidis...

group W-135; zwitterionization to convert T-independent into T-dependent immunogenic bacterial capsular saccharides for use as vaccines

Neisseria meningitidis...

group Y; zwitterionization to convert T-independent into T-dependent immunogenic bacterial capsular saccharides for use as vaccines Streptococcus group B...

Ia, Ib, II, III and V; zwitterionization to convert T-independent into T-dependent immunogenic bacterial capsular saccharides for use as vaccines

Acetyl group...

N-; zwitterionization to convert T-independent into T-dependent immunogenic bacterial capsular saccharides for use as vaccines Functional groups...

neutral, anionic and cationic; zwitterionization to convert T-independent into T-dependent immunogenic bacterial capsular saccharides for use as vaccines
Physical and chemical properties...

pKb; zwitterionization to convert T-independent into T-dependent

immunogenic bacterial capsular saccharides for use as vaccines
Amines, biological studies...

secondary; zwitterionization to convert T-independent into T-dependent

immunogenic bacterial capsular saccharides for use as vaccines Cell activation... T cell; zwitterionization to convert T-independent into T-dependent

immunogenic bacterial capsular saccharides for use as vaccines Carbohydrates, biological studies... Polysaccharides, biological studies...

Oligosaccharides, biological studies...
zwitterionic; zwitterionization to convert T-independent into

T-dependent immunogenic bacterial capsular saccharides for use as vaccines

Eubacteria.. Antigens.. Zwitterions.. Carboxyl group.. Amino group... Streptococcus agalactiae.. Monosaccharides.. Neisseria meningitidis.. Aldehydes, biological studies... Formyl group... Streptococcus pneumoniae... Bacteroides fragilis... Vaccines... T cell(lymphocyte)... zwitterionization to convert T-independent into T-dependent immunogenic

zwitterionization to convert T-independent into T-dependent immunogenic bacterial capsular saccharides for use as vaccines CAS REGISTRY NUMBERS:

50-00-0 127-17-3 64-19-7 24959-67-9 59-23-4 10028-15-6 10102-43-9

```
biological studies, zwitterionization to convert T-independent into T-dependent immunogenic bacterial capsular saccharides for use as vaccines
```

131-48-6 7512-17-6 moiety; zwitterionization to convert T-independent into T-dependent immunogenic bacterial capsular saccharides for use as vaccines

927927-51-3 927927-52-4 927927-53-5 927927-54-6 927927-55-7 927927-56-8 927927-56-9 927927-56-9 927927-56-9 927927-66-4 927927-66-6 927927-66-6 927927-66-7 927927-66-8 927927-65-9 927927-66-0 927927-66-9 927927-66-0 927927-66-0 927927-68-2 unclaimed sequence; zwitterionization to convert T-independent into T-dependent immunogenic

bacterial capsular saccharides for use as vaccines
927927-69-3 Unclaimed; zwitterionization to convert T-independent into
T-dependent immunogenic bacterial capsular saccharides for use as

vaccines
7790-28-5 151-51-9 2564-83-2 14380-61-1 52720-51-1 9028-79-9
zwitterionization to convert T-independent into T-dependent immunogenic
bacterial capsular saccharides for use as vaccines

10/7/35 (Item 14 from file: 399) DIALOG(R)File 399:CA SEARCH(R)

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```
145487661 CA: 145(25)487661z PATENT
```

Preparation of oligosaccharides conjugated with proteins or toxins as meningitidis A vaccines

INVENTOR(AUTHOR): Oscarson, Stefan; Teodorovic, Peter; Costantino, Paolo LOCATION: Italy

ASSIGNEE: Novartis Vaccines and Diagnostics S.r.l.; Stockholm University PATENT: PCT International; WO 2006120576 A2 DATE: 20061116

APPLICATION: WO 2006IB1703 (20060508) *US 2005PV678289 (20050506) PAGES: 87pp. CODEN: PIXXD2 LANGUAGE: English

PATENT CLASSIFICATIONS:

IPCR/8 + Level Value Position Status Version Action Source Office:

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C07H-0011/04
            A I F B 20060101
                                       H EP
A61K-0039/095
             A I L B 20060101
                                        H EP
A61P-0031/04
             A I L B 20060101
                                       H EP
C07H-0007/00
             A I L B 20060101
                                       H EP
C07H-0013/00
             A I L B 20060101
                                        H EP
C07H-0015/04
             A I L B 20060101
            A I L B 20060101
A61K-0031/702
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DESIGNATED COUNTRIES: AB; AG; AL; AM; AT; AU; AD; BA; BB; BG; BR; BW; BY; BZ; CA; CH; CN; CO; CR; CU; CZ; DE; DK; DM; DZ; EC; EE; EG; ES; FI; GB; GD; GE; GH; GN; HR; HU; ID; II; IN; IS; JF; KE; KG; KN; KN; KP; KR; KZ; LC; LK; LI; LI]; LU; LV; LY; MA; MD; MG; MK; MN; MM; MX; MZ; NA; NG; NI; NO; NZ; OM; PG; PH; PL; PT; RO; RU; SC; SD; SE; SG; SK; SL; SM; SY; TJ; TM; TN; TT; TZ; UA; UG; US; UZ; VC; VN; YU; ZA DESIGNATED REGIONAL: AT; BE; BG; CH; CY; CZ; DE; DK; EE; ES; FI; FR; GB; GR; HU; IE; IS; IT; LT; LU; LV; MC; NI; PL; PT; RO; SE; SI; SK; TR; BF; BJ; CF; GG; C1; CM; GA; GN; GQ; GW; ML; MR; NE; SN; TD; TG; BN; GH; GM; KE; LS; MW; MZ; NA; SD; SL; SZ; TZ; UG; ZM; ZM; AM; AZ; BY; KG; KZ; MD; RU; TJ; TM

CA215002 Immunochemistry

CA204XXX Toxicology

CA263XXX Pharmaceuticals

IDENTIFIERS: Neisseria meningitidis oligosaccharide protein toxin conjugate meningitis A vaccine

DESCRIPTORS:

SECTION:

Immunostimulants...

adjuvants; preparation of oligosaccharides conjugated with proteins or toxins as meningitidis ${\tt A}$ vaccines

Functional groups...

alkoxy groups; preparation of oligosaccharides conjugated with proteins or toxins as meningitidis A vaccines

Functional groups...

alkoxycarbonyl groups; preparation of oligosaccharides conjugated with proteins or toxins as meningitidis A vaccines

Streptococcus pneumoniae...

antigen; preparation of oligosaccharides conjugated with proteins or toxins as meningitidis A vaccines

Functional groups ... azido group; preparation of oligosaccharides conjugated with proteins or toxins as meningitidis A vaccines

Toxins...

bacterial; preparation of oligosaccharides conjugated with proteins or toxins as meningitidis A vaccines

Reagents..

bifunctional; preparation of oligosaccharides conjugated with proteins or toxins as meningitidis A vaccines

Oligosaccharides, biological studies... Proteins... Antigens... Toxins...

Polysaccharides, biological studies...

conjugates; preparation of oligosaccharides conjugated with proteins or toxins as meningitidis A vaccines Carboxvlic acids, biological studies ...

dicarboxylic: preparation of oligosaccharides conjugated with proteins or toxins as meningitidis A vaccines

Toxins... diphtheria, conjugates; preparation of oligosaccharides conjugated with proteins or toxins as meningitidis A vaccines

Toxoids...

diphtheria; preparation of oligosaccharides conjugated with proteins or toxins as meningitidis A vaccines

Neisseria meningitidis...

group A; preparation of oligosaccharides conjugated with proteins or toxins as meningitidis A vaccines

Neisseria meningitidis...

group B; preparation of oligosaccharides conjugated with proteins or toxins as meningitidis A vaccines Neisseria meningitidis...

group C; preparation of oligosaccharides conjugated with proteins or toxins as meningitidis A vaccines Neisseria meningitidis...

group W-135; preparation of oligosaccharides conjugated with proteins or toxins as meningitidis A vaccines

Neisseria meningitidis...

group Y; preparation of oligosaccharides conjugated with proteins or toxins as meningitidis A vaccines Toxins...

heat-labile; preparation of oligosaccharides conjugated with proteins or toxins as meningitidis A vaccines

Reaction...

Mitsunobu; preparation of oligosaccharides conjugated with proteins or toxins as meningitidis A vaccines

Toxins...

pertussis; preparation of oligosaccharides conjugated with proteins or toxins as meningitidis A vaccines

Functional groups...

phosphonate group; preparation of oligosaccharides conjugated with proteins or toxins as meningitidis A vaccines

Oligosaccharides, biological studies... Proteins... Antigens... Escherichia coli... Pseudomonas aeruginosa... Exotoxins... Hydroxyl group... Amino

group... Acyl groups... Protective groups... Amide group...

Carbohydrates, biological studies... Vaccines... Meningitis... Drug delivery

```
systems... Electrophiles... Nucleophiles... Antibodies and Immunoglobulins
... Albumins, biological studies... Polysaccharides, biological studies...
Alums...
   preparation of oligosaccharides conjugated with proteins or toxins as
   meningitidis A vaccines
   T; preparation of oligosaccharides conjugated with proteins or toxins as
   meningitidis A vaccines
Toxins...
   tetanus; preparation of oligosaccharides conjugated with proteins or toxins
   as meningitidis A vaccines
Eubacteria...
   toxin; preparation of oligosaccharides conjugated with proteins or toxins as
   meningitidis A vaccines
 CAS REGISTRY NUMBERS:
3458-28-4D -containing oligosaccharides, preparation of oligosaccharides conjugated
   with proteins or toxins as meningitidis A vaccines
124-04-9 biological studies, preparation of oligosaccharides conjugated with
   proteins or toxins as meningitidis A vaccines
172223-08-4 172223-09-5P 914641-02-4P 914641-03-5P 914641-04-6P
   914641-05-7P 914641-06-8P 914641-07-9 497096-20-5 914641-08-0P
   914641-09-1P 914641-10-4P 914641-11-5P 914641-12-6P 914641-13-7P
   914641-14-8P 68733-20-0 402831-54-3P 870073-99-7P 870074-00-3P
   870074-01-4P 870074-02-5P 870074-04-7P 870074-06-9P 870074-08-1P
   870074-12-7P 870074-14-9P 870074-16-1P 870074-18-3P 497096-09-0P
   870074-19-4P 870074-20-7P 870074-21-8P 870074-23-0P 870074-25-2P
   870074-27-4P 914641-15-9P 914641-17-1P 870074-31-0P 600173-37-3
   505-48-6 2892-51-5 preparation of oligosaccharides conjugated with
   proteins or toxins as meningitidis A vaccines
914641-18-2DP protein conjugates, preparation of oligosaccharides conjugated
   with proteins or toxins as meningitidis A vaccines
10/7/36
            (Item 15 from file: 399)
DIALOG(R) File 399:CA SEARCH(R)
(c) 2011 American Chemical Society. All rts. reserv.
 145209382
             CA: 145(11)209382h
 Conjugation of streptococcal capsular saccharides
 INVENTOR(AUTHOR): Berti, Francesco
 LOCATION: Italy
 ASSIGNEE: Chiron Srl
 PATENT: PCT International ; WO 200682530 A2 DATE: 20060810
 APPLICATION: WO 2006IB756 (20060201) *GB 20052095 (20050201)
 PAGES: 48pp. CODEN: PIXXD2 LANGUAGE: English
 PATENT CLASSIFICATIONS:
   IPCR/8 + Level Value Position Status Version Action Source Office:
     A61K-0047/48 A I F B 20060101
 DESIGNATED COUNTRIES: AE; AG; AL; AM; AT; AU; AZ; BA; BB; BG; BR; BW; BY;
BZ; CA; CH; CN; CO; CR; CU; CZ; DE; DK; DM; DZ; EC; EE; EG; ES; FI; GB; GD;
GE; GH; GM; HR; HU; ID; IL; IN; IS; JP; KE; KG; KM; KN; KP; KR; KZ; LC; LK;
LR; LS; LT; LU; LV; LY; MA; MD; MG; MK; MN; MW; MX; MZ; NA; NG; NI; NO; NZ;
OM; PG; PH; PL; PT; RO; RU; SC; SD; SE; SG; SK; SL; SM; SY; TJ; TM; TN; TR;
TT; TZ; UA; UG; US; UZ; VC; VN; YU; ZA DESIGNATED REGIONAL: AT; BE; BG; CH
; CY; CZ; DE; DK; EE; ES; FI; FR; GB; GR; HU; IE; IS; IT; LT; LU; LV; MC;
NL; PL; PT; RO; SE; SI; SK; TR; BF; BJ; CF; CG; CI; CM; GA; GN; GQ; GW; ML;
MR; NE; SN; TD; TG; BW; GH; GM; KE; LS; MW; MZ; NA; SD; SL; SZ; TZ; UG; ZM;
ZW; AM; AZ; BY; KG; KZ; MD; RU; TJ; TM
 SECTION:
   CA216002 Fermentation and Bioindustrial Chemistry
   CA233XXX Carbohydrates
```

CA244XXX Industrial Carbohydrates

```
IDENTIFIERS: streptococcal capsular saccharide conjugation
  DESCRIPTORS:
Polysaccharides, reactions...
    capsular; conjugation of streptococcal capsular saccharides
Streptococcus agalactiae... Capsule (microbial)... Oxidation... Linking
agents... Sialic acids... Human... Reducing agents... Vaccines...
    conjugation of streptococcal capsular saccharides
Proteins...
    D, as a carrier mol.; conjugation of streptococcal capsular saccharides
Toxoids...
    diphtheria, as a carrier mol.; conjugation of streptococcal capsular
    saccharides
Antigens...
    epitope; conjugation of streptococcal capsular saccharides
Proteins..
    from Streptococcus agalactiae, as a carrier mol.; conjugation of
    streptococcal capsular saccharides
Amines, reactions...
    primary; conjugation of streptococcal capsular saccharides
Amination...
    reductive; conjugation of streptococcal capsular saccharides
Albumins, reactions...
    serum, human, as a carrier mol.; conjugation of streptococcal capsular
    saccharides
Proteins...
    synthetic, multiple CD4+ epitopes, as a carrier mol.; conjugation of
    streptococcal capsular saccharides
Toxoids...
   tetanus, as a carrier mol.; conjugation of streptococcal capsular
    saccharides
 CAS REGISTRY NUMBERS:
7790-28-5 6066-82-6 59156-70-6 conjugation of streptococcal capsular
    saccharides
7664-41-7 processes, conjugation of streptococcal capsular saccharides
14798-03-9D salts, conjugation of streptococcal capsular saccharides
15056-35-6D salts, salts; conjugation of streptococcal capsular
    saccharides
 10/7/37
             (Item 16 from file: 399)
DIALOG(R)File 399:CA SEARCH(R)
(c) 2011 American Chemical Society. All rts. reserv.
              CA: 145(11)209376j
  145209376
                                   PATENT
  Purification of streptococcal capsular polysaccharide
  INVENTOR (AUTHOR): Costantino, Paolo
  LOCATION: Italy
  ASSIGNEE: Chiron Srl
  PATENT: PCT International ; WO 200682527 A2 DATE: 20060810
  APPLICATION: WO 2006IB626 (20060201) *GB 20052096 (20050201)
  PAGES: 39pp. CODEN: PIXXD2 LANGUAGE: English
  PATENT CLASSIFICATIONS:
    IPCR/8 + Level Value Position Status Version Action Source Office:
                       A I F B 20060101
A I L B 20060101
      C08B-0037/00
                                                        H EP
      C12P-0019/04
  DESIGNATED COUNTRIES: AE; AG; AL; AM; AT; AU; AZ; BA; BB; BG; BR; BW; BY;
BZ; CA; CH; CN; CO; CR; CU; CZ; DE; DK; DM; DZ; EC; EE; EG; ES; FI; GB; GD;
GE; GH; GM; HR; HU; ID; IL; IN; IS; JP; KE; KG; KM; KN; KP; KR; KZ; LC; LK;
LR; LS; LT; LU; LV; LY; MA; MD; MG; MK; MN; MW; MX; MZ; NA; NG; NI; NO; NZ;
OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SM, SY, TJ, TM, TN, TR,
TT; TZ; UA; UG; US; UZ; VC; VN; YU; ZA DESIGNATED REGIONAL: AT; BE; BG; CH
; CY; CZ; DE; DK; EE; ES; FI; FR; GB; GR; HU; IE; IS; IT; LT; LU; LV; MC;
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NL; PL; PT; RO; SE; SI; SK; TR; BF; BJ; CF; CG; CI; CM; GA; GN; GQ; GW; ML;
MR; NE; SN; TD; TG; BW; GH; GM; KE; LS; MW; MZ; NA; SD; SL; SZ; TZ; UG; ZM;
ZW; AM; AZ; BY; KG; KZ; MD; RU; TJ; TM
 SECTION:
    CA216001 Fermentation and Bioindustrial Chemistry
   CA233XXX Carbohydrates
    CA244XXX Industrial Carbohydrates
  IDENTIFIERS: streptococcal capsular polysaccharide purifn
  DESCRIPTORS:
Extraction...
    base; purification of streptococcal capsular polysaccharide
Polysaccharides, preparation...
    capsular; purification of streptococcal capsular polysaccharide
Detergents...
   cationic; purification of streptococcal capsular polysaccharide
Hydrolysis..
    chemical; purification of streptococcal capsular polysaccharide
Hydrolysis...
    enzymic; purification of streptococcal capsular polysaccharide
Filtration...
    microfiltration; purification of streptococcal capsular polysaccharide
Streptococcus agalactiae... Capsule (microbial)... Cations...
Alcohols, processes... Precipitation (chemical)... Separation...
Centrifugation... Fermentation... Ultrafiltration... Solubilization...
Proteins... Nucleic acids... Vaccines...
    purification of streptococcal capsular polysaccharide
Filtration...
   tangential-flow filtration; purification of streptococcal capsular
    polysaccharide
  CAS REGISTRY NUMBERS:
64-17-5 67-63-0 16397-91-4 22537-22-0 14127-61-8 10043-52-4
    processes, purification of streptococcal capsular polysaccharide
55466-22-3 9012-33-3 9025-82-5 57-09-0 10549-76-5 purification of
    streptococcal capsular polysaccharide
 10/7/38
             (Item 17 from file: 399)
DIALOG(R)File 399:CA SEARCH(R)
(c) 2011 American Chemical Society. All rts. reserv.
              CA: 143(1)5815a
                                 JOURNAL
  Size determination of bacterial capsular oligosaccharides used to prepare
  conjugate vaccines against Neisseria meningitidis groups Y and W135
 AUTHOR(S): Bardotti, Angela; Averani, Giovanni; Berti, Francesco; Berti,
Stefania; Galli, Chiara; Giannini, Sara; Fabbri, Barbara; Proietti, Daniela
; Ravenscroft, Neil; Ricci, Stefano
  LOCATION: Chiron Vaccines, I-53100, Siena, Italy
```

LOCATION: Chiron Vaccines, I-53100, Siena, Italy
JOURNAL: Vaccine | DATE: 2005 VOLUME: 23 NUMBER: 16 PAGES:
1887-1899 CODEN: VACCDE ISSN: 0264-410X PUBLISHER ITEM IDENTIFIER:
0264-410X(04)00790-X LANGUAGE: English PUBLISHER: Elsevier B.V.
SECTION:
CA215002 Immunochemistry
IDENTIFIERS: size detn bacteria capsule oligosaccharide Neisseria vaccine
DESCRIPTORS:
Polymerization...

average degree of; size determination of bacterial capsular oligosaccharides used to prepare conjugate vaccines against Neisseria meningitidis groups Y and

W135

Oligosaccharides, biological studies... Vaccines... Neisseria meningitidis ... Anion exchange chromatography...

size determination of bacterial capsular oligosaccharides used to prepare

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(Item 18 from file: 399)
 10/7/39
DIALOG(R)File 399:CA SEARCH(R)
(c) 2011 American Chemical Society. All rts. reserv.
              CA: 142(25)461845a
                                    JOURNAL
 The concept of "tailor-made", protein-based, outer membrane vesicle
  vaccines against meningococcal disease
 AUTHOR(S): Holst, Johan; Feiring, Berit; Naess, Lisbeth M.; Norheim,
Gunnstein; Kristiansen, Paul; Hoiby, E. Arne; Bryn, Klaus; Oster, Philipp;
Costantino, Paolo; Taha, Muhamed-Kheir; Alonso, Jean-Michel; Caugant,
Dominique A.; Wedege, Elisabeth; Aaberge, Ingeborg S.; Rappuoli, Rino;
Rosenqvist, Einar
 LOCATION: Norwegian Institute of Public Health, Oslo, Norway
 JOURNAL: Vaccine (Vaccine) DATE: 2005 VOLUME: 23 NUMBER: 17-18
  PAGES: 2202-2205 CODEN: VACCDE ISSN: 0264-410X
  PUBLISHER ITEM IDENTIFIER: 0264-410X(05)00063-0 LANGUAGE: English
  PUBLISHER: Elsevier B.V.
  SECTION:
   CA215002 Immunochemistry
  IDENTIFIERS: Neisseria meningitidis vaccine outer membrane vesicle
  DESCRIPTORS:
Infection...
    bacterial; concept of tailor-made protein-based outer membrane vesicle
    vaccines against meningococcal disease
Neisseria meningitidis... Vaccines... Human...
    concept of tailor-made protein-based outer membrane vesicle vaccines
   against meningococcal disease
Cell wall...
   outer membrane; concept of tailor-made protein-based outer membrane
    vesicle vaccines against meningococcal disease
Organelle...
   vesicle; concept of tailor-made protein-based outer membrane vesicle
   vaccines against meningococcal disease
 10/7/40
             (Item 19 from file: 399)
DIALOG(R)File 399:CA SEARCH(R)
(c) 2011 American Chemical Society. All rts. reserv.
              CA: 142(20)372474m
                                   PATENT
  Acetylated meningococcal capsular oligosaccharides
  INVENTOR(AUTHOR): Costantino, Paolo
  LOCATION: Italy
  ASSIGNEE: Chiron S.r.l.
  PATENT: PCT International ; WO 200533148 A1 DATE: 20050414
 APPLICATION: WO 2004IB3366 (20041004) *GB 200323103 (20031002)
  PAGES: 42 pp. CODEN: PIXXD2 LANGUAGE: English
  PATENT CLASSIFICATIONS:
    CLASS: C08B-037/00A; A61K-039/095B; A61K-031/715B
  DESIGNATED COUNTRIES: AE; AG; AL; AM; AT; AU; AZ; BA; BB; BG; BR; BW; BY;
BZ; CA; CH; CN; CO; CR; CU; CZ; DE; DK; DM; DZ; EC; EE; EG; ES; FI; GB; GD;
GE; GH; GM; HR; HU; ID; IL; IN; IS; JP; KE; KG; KP; KR; KZ; LC; LK; LR; LS;
LT; LU; LV; MA; MD; MG; MK; MN; MW; MX; MZ; NA; NI; NO; NZ; OM; PG; PH; PL;
PT; RO; RU; SC; SD; SE; SG; SK; SL; SY; TJ; TM; TN; TR; TT; TZ; UA; UG; US;
UZ; VC; VN; YU; ZA; ZM; ZW DESIGNATED REGIONAL: BW; GH; GM; KE; LS; MW; MZ
; NA; SD; SL; SZ; TZ; UG; ZM; ZW; AM; AZ; BY; KG; KZ; MD; RU; TJ; TM; AT;
BE; BG; CH; CY; CZ; DE; DK; EE; ES; FI; FR; GB; GR; HU; IE; IT; LU; MC; NL;
PL; PT; RO; SE; SI; SK; TR; BF; BJ; CF; CG; CI; CM; GA; GN; GQ; GW; ML; MR;
NE; SN; TD; TG
```

SECTION:

CA215002 Immunochemistry

IDENTIFIERS: acetylated capsular polysaccharide Neisseria vaccine DESCRIPTORS:

Human...

acetylated sialate oligosaccharides of serogroups W135 and Y of Neisseria meningitidis for vaccination

Polysaccharides, biological studies...

capsular, fragments, conjugates; for vaccination against serogroups W135 and Y of Neisseria meningitidis

Proteins...

D, conjugates with acetylated sialate oligosaccharides; for vaccination against serogroups W135 and Y of Neisseria meningitidis

Toxoids...

diphtheria, conjugates with acetylated sialate oligosaccharides; for vaccination against serogroups W135 and Y of Neisseria meningitidis...

group A; vaccination with acetylated sialate oligosaccharides of serogroups W135 and Y of N. meningitidis and saccharide antigen from Neisseria meningitidis...

group W-135; acetylated sialate oligosaccharide conjugates for vaccination against

Neisseria meningitidis...

group Y; acetylated sialate oligosaccharide conjugates for vaccination against

Antigens...

microbial; in combination vaccination with acetylated sialate oligosaccharides of serogroups W135 and Y of Neisseria meningitidis Vaccines...

of acetylated sialate oligosaccharides of serogroups W135 and Y of Neisseria meningitidis

Acetyl group...

of Neisseria meningitidis capsular polysaccharide affects immunogenicity

Toxoids...

tetanus, conjugates with acetylated sialate oligosaccharides; for vaccination against serogroups W135 and Y of Neisseria meningitidis Streptococcus pneumoniae... Hepatitis A virus... Hepatitis B virus... Bordetella pertussis... Human poliovirus...

vaccination with acetylated sialate oligosaccharides of serogroups W135 and Y of Neisseria meningitidis and antigen from

Antibodies and Immunoglobulins...

vaccination with acetylated sialate oligosaccharides of serogroups W135 and Y of Neisseria meningitidis for elicitation of Meningitis...

vaccination with acetylated sialate oligosaccharides of serogroups W135 and Y of Neisseria meningitidis for protection against

Sialooligosaccharides...

7-acetylated sialate- or 9-acetylated sialate-containing, conjugates; for vaccination against serogroups W135 and Y of Neisseria meningitidis CAS REGISTRY NUMBERS:

736884-44-9DP 849592-59-2DP acetyl derivs., carrier protein conjugates, repeating unit; acetylated sialate oligosaccharides of serogroups W135 and Y of Neisseria menincitidis for vaccination

600173-37-3D conjugates with acetylated sialate oligosaccharides, for vaccination against serogroups W135 and Y of Neisseria meningitidis

10/7/41 (Item 20 from file: 399)

DIALOG(R)File 399:CA SEARCH(R)

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141189619
              CA: 141(12)189619c
                                    PATENT
  Injectable immunogenic composition comprising capsular saccharides from
 Neisseria meningitidis B, A, C, W135 and Y as vaccines against multiple
 meningococcal serogroups
  INVENTOR (AUTHOR): Costantino, Paolo
  LOCATION: Italy
  ASSIGNEE: Chiron SRL
  PATENT: PCT International; WO 200467030 A2 DATE: 20040812
 APPLICATION: WO 2004IB651 (20040130) *GB 20032217 (20030130) *GB
200323101 (20031002)
  PAGES: 46 pp. CODEN: PIXXD2 LANGUAGE: English
  PATENT CLASSIFICATIONS:
    CLASS: A61K-039/095A; A61K-039/39B; C07K-014/22B; A61K-039/02B;
A61K-039/09B
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MOIA-U3/U3P DESIGNATED COUNTRIES: AE; AE; AE; AL; AL; AM; AM; AM; AT; AT; AU; AZ; AZ; BA; BB; BG; BG; BR; BR; BW; BY; BY; BZ; BZ; CA; CH; CN; CN; CO; CO; CR; CR; CC; CU; CZ; CZ; DE; DE; DK; DK; DM; DZ; EC; EC; EE; EE; EG; ES; ES; FI; GB; GB; GB; GB; GB; GH; GM; HB; HB; HB; HU; HU; ID; IL; IN; IS; JP; JP; KE; KE; KG; KG; KP; KP; KP; KR; KR; KZ; KZ; KZ; KC; KE; KF; LS; LS; LT; LU; LV; MA; MD; MG; MK; MN; MW; MX; MX; MZ; MZ; NA; NI
SECTION:

CA215002 Immunochemistry CA263XXX Pharmaceuticals

IDENTIFIERS: Neisseria meningitidis A B C W135 Y capsular saccharide, Haemophilus influenzae Streptococcus pneumoniae Neisseria meningitidis meningococcal vaccine injection DESCRIPTORS:

Immunostimulants...

adjuvants; injectable immunogenic composition comprising capsular saccharides from Neisseria meningitidis B, A, C, W135 and Y as vaccines against bacterial meningitis
Meningitis...

bacterial; injectable immunogenic composition comprising capsular saccharides from Neisseria meningitidis B, A, C, W135 and Y as vaccines against bacterial meningitis

Polysaccharides, biological studies...

capsular; injectable immunogenic composition comprising capsular saccharides from Neisseria meningitidis B, A, C, W135 and Y as vaccines against bacterial meningitis

Drug delivery systems...

carriers, injectable immunogenic composition comprising capsular saccharides from Neisseria meningitidis B, A, C, W135 and Y as vaccines against bacterial meningitis

Proteins... Antigens...

conjugates; injectable immunogenic composition comprising capsular saccharides from Neisseria meningitidis B, A, C, W135 and Y as vaccines against bacterial meningitis

Proteins...

D; injectable immunogenic composition comprising capsular saccharides from Neisseria meningitidis B, A, C, W135 and Y as vaccines against bacterial meningitis

Toxins...

diphtheria, conjugates; injectable immunogenic composition comprising capsular saccharides from Neisseria meningitidis B, A, C, W135 and Y as vaccines against bacterial meningitis

Toxoids...

diphtheria; injectable immunogenic composition comprising capsular saccharides from Neisseria meningitidis B, A, C, W135 and Y as vaccines against bacterial meningitis

Drug delivery systems...

freeze-dried; injectable immunogenic composition comprising capsular saccharides from Neisseria meningitidis B, A, C, W135 and Y as vaccines

against bacterial meningitis

Neisseria meningitidis...

group A; injectable immunogenic composition comprising capsular saccharides from Neisseria meningitidis B, A, C, W135 and Y as vaccines against bacterial meningitis

Neisseria meningitidis...

group B; injectable immunogenic composition comprising capsular saccharides from Neisseria meningitidis B, A, C, W135 and Y as vaccines against bacterial meningitis

Neisseria meningitidis...

group C; injectable immunogenic composition comprising capsular saccharides from Neisseria meningitidis B, A, C, W135 and Y as vaccines against bacterial meningitis

Neisseria meningitidis...

group W-135; injectable immunogenic composition comprising capsular saccharides from Neisseria meningitidis B, A, C, W135 and Y as vaccines against bacterial meningitis

Neisseria meningitidis...

group Y; injectable immunogenic composition comprising capsular saccharides from Neisseria meningitidis B, A, C, W135 and Y as vaccines against bacterial meningitis

Vaccines.. Eubacteria... Neisseria meningitidie... Streptococcus pneumoniae... Oligosaccharides, biological studies... Hydroxyl group... Protective groups... Antibodies and Immunoglobulins... Antibacterial agents ... Fusion proteins(chimeric proteins)... Test kits... Protein sequences... Human...

injectable immunogenic composition comprising capsular saccharides from Neisseria meningitidis B, A, C, W135 and Y as vaccines against bacterial meningitis

Drug delivery systems...

injections, i.m.; injectable immunogenic composition comprising capsular saccharides from Neisseria meningitidis B, A, C, W135 and Y as vaccines against bacterial meningitis

Drug delivery systems...

injections, s.c.; injectable immunogenic composition comprising capsular saccharides from Neisseria meningitidis B, A, C, W135 and Y as vaccines against bacterial meningitis

Drug delivery systems ...

injections; injectable immunogenic composition comprising capsular saccharides from Neisseria meningitidis B, A, C, W135 and Y as vaccines against bacterial meningitis

Drug delivery systems...

ligs.; injectable immunogenic composition comprising capsular saccharides from Neisseria meningitidis B, A, C, W135 and Y as vaccines against bacterial meningitis

Antigens...

LytA; injectable immunogenic composition comprising capsular saccharides from Neisseria meninguitidis B, A, C, W135 and Y as vaccines against bacterial meninguitis

Antigens...

LytB; injectable immunogenic composition comprising capsular saccharides from Neisseria meningitidis B, A, C, W135 and Y as vaccines against bacterial meningitis

Antigens...

LytC; injectable immunogenic composition comprising capsular saccharides from Neisseria meningitidis B, A, C, W135 and Y as vaccines against bacterial meningitis

Antigens...

Nada (Neisserial adhesin A); injectable immunogenic composition comprising capsular saccharides from Neisseria meningitidis B, A, C, W135 and Y as vaccines against bacterial meningitis

Antigens...

phtA; injectable immunogenic composition comprising capsular saccharides from Neisseria meningitidis B, A, C, W135 and Y as vaccines against bacterial meningitis

Antigens...

phtB; injectable immunogenic composition comprising capsular saccharides from Neisseria meningitidis B, A, C, W135 and Y as vaccines against bacterial meningitis

Antigens..

phtD; injectable immunogenic composition comprising capsular saccharides from Neisseria meningitidis B, A, C, W135 and Y as vaccines against bacterial meningitis

Antigens...

phtE; injectable immunogenic composition comprising capsular saccharides from Neisseria meningitidis B, A, C, W135 and Y as vaccines against bacterial meningitis

Antigens...

SpsA; injectable immunogenic composition comprising capsular saccharides from Neisseria meningitidis B, A, C, Wl35 and Y as vaccines against bacterial meningitis

Antigens...

Sp101; injectable immunogenic composition comprising capsular saccharides from Neisseria meningitidis B, A, C, W135 and Y as vaccines against bacterial meningitis

Antigens...

Sp125; injectable immunogenic composition comprising capsular saccharides from Neisseria meningitidis B, A, C, W135 and Y as vaccines against bacterial meningitis

Antigens...

Sp128; injectable immunogenic composition comprising capsular saccharides from Neisseria meningitidis B, A, C, W135 and Y as vaccines against bacterial meningitis

Antigens...

Sp130; injectable immunogenic composition comprising capsular saccharides from Neisseria meningitidis B, A, C, W135 and Y as vaccines against bacterial meningitis

Toxoids...

tetanus; injectable immunogenic composition comprising capsular saccharides from Neisseria meningitidis B, A, C, W135 and Y as vaccines against bacterial meningitis

Haemophilus influenzae...

type b; injectable immunogenic composition comprising capsular saccharides from Neisseria meningitidis B, A, C, W135 and Y as vaccines against bacterial meningitis

Antigens...

287; injectable immunogenic composition comprising capsular saccharides from Neisseria meningitidis B, A, C, W135 and Y as vaccines against bacterial meningities

Antigens...

741; injectable immunogenic composition comprising capsular saccharides from Neisseria meningitidis B, A, C, W135 and Y as vaccines against bacterial meningitis

Antigens...

936; injectable immunogenic composition comprising capsular saccharides from Neisseria meningitidis B, A, C, W135 and Y as vaccines against bacterial meningitis

Antigens...

953; injectable immunogenic composition comprising capsular saccharides from Neisseria meningitidis B, A, C, W135 and Y as vaccines against bacterial meningitis

CAS REGISTRY NUMBERS:

738308-30-0P 738308-31-1P 738308-32-2P 738308-33-3P 738308-34-4P 738308-35-5P 738308-36-6P 738308-37-7P 738308-38-8P 738308-39-9P

- 738308-40-2P amino acid sequence; injectable immunogenic composition comprising capsular saccharides from Neisseria meningitidis B, A, C, W135 and Y as vaccines against bacterial meningitis
- 7429-90-5D salts, injectable immunogenic composition comprising capsular saccharides from Neisseria meningitidis B, A, C, W135 and Y as vaccines against bacterial meningitis
- 219724-66-0 unclaimed sequence; injectable immunogenic composition comprising capsular saccharides from Neisseria meningitidis B, A, C, W135 and Y as vaccines against multiple meningococcal serogroups

10/7/42 (Item 21 from file: 399)

DIALOG(R)File 399:CA SEARCH(R)

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CA: 141(4)52579v JOURNAL

Modulation of immune response to group C meningococcal conjugate vaccine given intranasally to mice together with the LTK63 mucosal adjuvant and the trimethyl chitosan delivery system

AUTHOR(S): Baudner, Barbara C.; Morandi, Maurizio; Giuliani, Marzia M.; Verhoef, J. Coos; Junginger, Hans E.; Costantino, Paolo; Rappuoli, Rino; Del Giudice, Giuseppe

LOCATION: Research Center, Chiron Srl, 53100, Siena, Italy

JOURNAL: J. Infect. Dis. (Journal of Infectious Diseases) DATE: 2004 VOLUME: 189 NUMBER: 5 PAGES: 828-832 CODEN: JIDIAQ ISSN: 0022-1899 LANGUAGE: English PUBLISHER: University of Chicago Press SECTION:

CA215003 Immunochemistry

IDENTIFIERS: vaccine Neisseria LTK63 adjuvant trimethyl chitosan delivery system DESCRIPTORS:

Immunostimulants...

adjuvants; modulation of immune response to group C meningococcal conjugate vaccine given intranasally to mice together with the LTK63 mucosal adjuvant and the tri-Me chitosan delivery system Vaccines...

conjugate; modulation of immune response to group C meningococcal conjugate vaccine given intranasally to mice together with the LTK63 mucosal adjuvant and the tri-Me chitosan delivery system Neisseria meningitidis...

group C; modulation of immune response to group C meningococcal conjugate vaccine given intranasally to mice together with the LTK63 mucosal adjuvant and the tri-Me chitosan delivery system

Antibodies and Immunoglobulins...

IgG; modulation of immune response to group C meningococcal conjugate vaccine given intranasally to mice together with the LTK63 mucosal adjuvant and the tri-Me chitosan delivery system

Drug delivery systems...

modulation of immune response to group C meningococcal conjugate vaccine given intranasally to mice together with the LTK63 mucosal adjuvant and the tri-Me chitosan delivery system

Immunization...

vaccination; modulation of immune response to group C meningococcal conjugate vaccine given intranasally to mice together with the LTK63 mucosal adjuvant and the tri-Me chitosan delivery system

CAS REGISTRY NUMBERS:

226416-68-8 modulation of immune response to group C meningococcal conjugate vaccine given intranasally to mice together with the LTK63 mucosal adjuvant and the tri-Me chitosan delivery system

9012-76-4D tri-Me derivs., modulation of immune response to group C meningococcal conjugate vaccine given intranasally to mice together with the LTK63 mucosal adjuvant and the tri-Me chitosan delivery system

```
Costantino, Paolo
  LOCATION: Italy
  ASSIGNEE: Chiron S.r.l.
  PATENT: PCT International; WO 200419992 A1 DATE: 20040311
  APPLICATION: WO 2003IB4194 (20030901) *GB 200220198 (20020830)
  PAGES: 30 pp. CODEN: PIXXD2 LANGUAGE: English
  PATENT CLASSIFICATIONS:
    CLASS: A61K-047/48A; C07H-013/12B; A61K-031/715B
  DESIGNATED COUNTRIES: AE; AG; AL; AM; AT; AU; AZ; BA; BB; BG; BR; BY; BZ;
CA; CH; CN; CO; CR; CU; CZ; DE; DK; DM; DZ; EC; EE; ES; FI; GB; GD; GE; GH;
GM; HR; HU; ID; IL; IN; IS; JP; KE; KG; KP; KR; KZ; LC; LK; LR; LS; LT; LU;
LV; MA; MD; MG; MK; MN; MW; MX; MZ; NI; NO; NZ; OM; PG; PH; PL; PT; RO; RU;
SC; SD; SE; SG; SK; SL; SY; TJ; TM; TN; TR; TT; TZ; UA; UG; US; UZ; VC; VN;
YU; ZA; ZM; ZW; AM; AZ; BY; KG; KZ; MD; RU DESIGNATED REGIONAL; GH; GM; KE
; LS; MW; MZ; SD; SL; SZ; TZ; UG; ZM; ZW; AT; BE; BG; CH; CY; CZ; DE; DK;
EE; ES; FI; FR; GB; GR; HU; IE; IT; LU; MC; NL; PT; RO; SE; SI; SK; TR; BF;
BJ; CF; CG; CI; CM; GA; GN; GQ; GW; ML; MR; NE; SN; TD; TG
 SECTION:
    CA263003 Pharmaceuticals
    CA215XXX Immunochemistry
   CA233XXX Carbohydrates
    CA234XXX Amino Acids, Peptides, and Proteins
  IDENTIFIERS: capsular oligosaccharide linker protein conjugate immunogen,
    Neisseria capsular polysaccharide protein conjugate meningitis
  DESCRIPTORS:
Immunostimulants...
    adjuvants; preparation of saccharide-protein conjugates with improved
    immunogenicity
Meningitis...
    bacterial; preparation of saccharide-protein conjugates with improved
    immunogenicity
Oligosaccharides, biological studies... Polysaccharides, biological studies
    capsular, conjugates with proteins; preparation of saccharide-protein
    conjugates with improved immunogenicity
Proteins... Toxins... Toxoids...
    conjugates with capsular saccharides; preparation of saccharide-protein
    conjugates with improved immunogenicity
Proteins...
    CRM197, conjugates with capsular saccharides; preparation of
    saccharide-protein conjugates with improved immunogenicity
Toxins... Toxoids...
    diphtheria, conjugates with capsular saccharides; preparation of
    saccharide-protein conjugates with improved immunogenicity
Neisseria meningitidis...
    group A; preparation of saccharide-protein conjugates with improved
    immunogenicity
Vaccines... Antigens...
```

preparation of saccharide-protein conjugates with improved immunogenicity

reductive; preparation of saccharide-protein conjugates with improved

PATENT

INVENTOR (AUTHOR): Giannozzi, Aldo; Averani, Giovanni; Norelli, Francesco;

(Item 22 from file: 399)

CA: 140(15)240996q

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Modified saccharides and their protein conjugates

DIALOG(R)File 399:CA SEARCH(R)

10/7/43

140240996

Amination...

immunogenicity
Drug delivery systems...

saccharide-protein conjugates with improved immunogenicity CAS REGISTRY NUMBERS:

530-62-1 41864-22-6 68985-05-7 102-09-0 506-68-3 75-44-5 32315-10-9 bifunctional reagent; preparation of saccharide-protein conjugates with improved immunogenicity

375345-20-3 preparation of saccharide-protein conjugates with improved immunogenicity

25895-60-7 reducing agent; preparation of saccharide-protein conjugates with improved immunogenicity

10/7/44 (Item 23 from file: 399) DIALOG(R)File 399:CA SEARCH(R)

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CA: 140(8)109759q JOURNAL

Water accessibility, aggregation, and motional features of polysaccharide-protein conjugate vaccines

AUTHOR(S): Berti, Francesco; Costantino, Paolo; Fragai, Marco; Luchinat, Claudio

LOCATION: IRIS, Chiron SPA, 53100, Siena, Italy

JOURNAL: Biophys. J. (Biophysical Journal) DATE: 2004 VOLUME: 86 NUMBER: 1, Pt. 1 PAGES: 3-9 CODEN: BIOJAU ISSN: 0006-3495 LANGUAGE: English PUBLISHER: Biophysical Society

SECTION: CA215002 Immunochemistry

IDENTIFIERS: polysaccharide protein conjugate vaccine water hydrodynamics DESCRIPTORS:

Vaccines...

conjugate; water accessibility, aggregation, and motional features of polysaccharide-protein conjugate vaccines

Aggregation... Hydration, chemical... Molecular dynamics...

Polysaccharides, biological studies... Proteins...

water accessibility, aggregation, and motional features of polysaccharide-protein conjugate vaccines

10/7/45 (Item 24 from file: 399)

DIALOG(R)File 399:CA SEARCH(R)

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139275732 CA: 139(18)275732z PATENT

Modified polysaccharide-protein conjugates having improved stability in water for use as vaccines

INVENTOR(AUTHOR): Costantino, Paolo; Berti, Francesco; Norelli, Francesco ; Bartoloni, Antonella

LOCATION: Italy

ASSIGNEE: Chiron S.r.l.

PATENT: PCT International ; WO 200380678 Al DATE: 20031002

APPLICATION: WO 2003IB1436 (20030326) *GB 20027117 (20020326) *GB

200220195 (20020830) *GB 200229494 (20021218) *GB 200230163 (20021224) PAGES: 58 pp. CODEN: PIXXD2 LANGUAGE: English

PATENT CLASSIFICATIONS:

CLASS: C08B-037/00A; A61K-039/385B; A61K-039/095B

DESIGNATED COUNTRIES: AE; AG; AL; AM; AT; AU; AZ; BA; BB; BG; BR; BY; BZ; CA; CH; CN; CO; CR; CU; CZ; DE; DK; DM; DZ; EC; EE; ES; FI; GB; GD; GE; GH; GM; HR; HU; ID; IL; IN; IS; JP; KE; KG; KP; KR; KZ; LC; LK; LR; LS; LT; LU; LV; MA; MD; MG; MK; MN; MW; MX; MZ; NI; NO; NZ; OM; PH; PL; PT; RO; RU; SC; SD; SE; SG; SK; SL; TJ; TM; TN; TR; TT; TZ; UA; UG; US; UZ; VC; VN; YU; ZA; ZM; ZW; AM; AZ; BY; KG; KZ; MD; RU; TJ; TM DESIGNATED REGIONAL: GH; GM; KE ; LS; MW; MZ; SD; SL; SZ; TZ; UG; ZM; ZW; AT; BE; BG; CH; CY; CZ; DE; DK; EE; ES; FI; FR; GB; GR; HU; IE; IT; LU; MC; NL; PT; RO; SE; SI; SK; TR; BF; BJ; CF; CG; CI; CM; GA; GN; GQ; GW; ML; MR; NE; SN; TD; TG

SECTION:

CA215002 Immunochemistry

CA209XXX Biochemical Methods

CA263XXX Pharmaceuticals

IDENTIFIERS: Neisseria meningitidis capsular saccharide oligosaccharide polysaccharide protein conjugate vaccine

DESCRIPTORS:

Immunostimulants...

adjuvants; modified polysaccharide-protein conjugates having improved stability in water for use as vaccines

Solvents...

aprotic; modified polysaccharide-protein conjugates having improved stability in water for use as vaccines

Toxins... Meningitis...

bacterial; modified polysaccharide-protein conjugates having improved stability in water for use as vaccines

Reagents... Crosslinking agents...

bifunctional; modified polysaccharide-protein conjugates having improved stability in water for use as vaccines

Functional groups ...

blocking; modified polysaccharide-protein conjugates having improved stability in water for use as vaccines

Drug delivery systems...

carriers; modified polysaccharide-protein conjugates having improved stability in water for use as vaccines

Proteins... Monosaccharides... Oligosaccharides, biological studies...

Polysaccharides, biological studies..

conjugates; modified polysaccharide-protein conjugates having improved stability in water for use as vaccines

Toxins...

diphtheria; modified polysaccharide-protein conjugates having improved stability in water for use as vaccines Functional groups ...

electron-withdrawing; modified polysaccharide-protein conjugates having improved stability in water for use as vaccines

enhancement; modified polysaccharide-protein conjugates having improved stability in water for use as vaccines

Neisseria meningitidis...

group A; modified polysaccharide-protein conjugates having improved stability in water for use as vaccines Neisseria meningitidis...

group C; modified polysaccharide-protein conjugates having improved stability in water for use as vaccines

Neisseria meningitidis...

group W-135; modified polysaccharide-protein conjugates having improved stability in water for use as vaccines

Neisseria meningitidis...

group Y; modified polysaccharide-protein conjugates having improved stability in water for use as vaccines

Glycosides...

linkage; modified polysaccharide-protein conjugates having improved stability in water for use as vaccines

Monosaccharides... Capsule (microbial)... Oligosaccharides, biological studies... Polysaccharides, biological studies... Hydroxyl group... Amino group... Bacteria (Eubacteria)... Vaccines... Antibodies... Mammalia...

Amines, biological studies... Glycoconjugates... modified polysaccharide-protein conjugates having improved stability in

water for use as vaccines Functional groups...

nitrogen-protecting; modified polysaccharide-protein conjugates having

improved stability in water for use as vaccines Solvents... organic; modified polysaccharide-protein conjugates having improved stability in water for use as vaccines Functional groups ... phosphodiester, bond; modified polysaccharide-protein conjugates having improved stability in water for use as vaccines Carbohydrates, biological studies... sugar phosphates; modified polysaccharide-protein conjugates having improved stability in water for use as vaccines CAS REGISTRY NUMBERS: 67-68-5 68-12-2 75-12-7 680-31-9 biological studies, modified polysaccharide-protein conjugates having improved stability in water for use as vaccines 1608-26-0 7226-23-5 127-19-5 530-62-1 41864-22-6 23814-12-2 102-09-0 506-68-3 75-44-5 32315-10-9 600173-37-3 7784-30-7 modified polysaccharide-protein conjugates having improved stability in water for use as vaccines (Item 25 from file: 399) 10/7/46 DIALOG(R)File 399:CA SEARCH(R) (c) 2011 American Chemical Society, All rts, reserv. CA: 138(9)121627m 138121627 Purification of bacterial capsular polysaccharide for use in combination vaccines INVENTOR (AUTHOR): Costantino, Paolo LOCATION: Italy ASSIGNEE: Chiron S.P.A. PATENT: PCT International ; WO 200307985 A2 DATE: 20030130 APPLICATION: WO 2002IB3191 (20020620) *GB 200115176 (20010620) PAGES: 49 pp. CODEN: PIXXD2 LANGUAGE: English PATENT CLASSIFICATIONS: CLASS: A61K-039/02A; A61K-039/095B; A61K-039/385B; A61P-031/04B DESIGNATED COUNTRIES: AE; AG; AL; AM; AT; AU; AZ; BA; BB; BG; BR; BY; BZ; CA; CH; CN; CO; CR; CU; CZ; DE; DK; DM; DZ; EC; EE; ES; FI; GB; GD; GE; GH; GM; HR; HU; ID; IL; IN; IS; JP; KE; KG; KP; KR; KZ; LC; LK; LR; LS; LT; LU;

LV; MA; MD; MG; MK; MN; MW; MX; MZ; NO; NZ; OM; PH; PL; PT; RO; RU; SD; SE; SG; SI; SK; SL; TJ; TM; TN; TR; TT; TZ; UA; UG; US; UZ; VN; YU; ZA; ZM; ZW; AM; AZ; BY; KG; KZ; MD; RU; TJ; TM DESIGNATED REGIONAL: GH; GM; KE; LS; MW ; MZ; SD; SL; SZ; TZ; UG; ZM; ZW; AT; BE; CH; CY; DE; DK; ES; FI; FR; GB; GR; IE; IT; LU; MC; NL; PT; SE; TR; BF; BJ; CF; CG; CI; CM; GA; GN; GQ; GW; ML; MR; NE; SN; TD; TG SECTION:

CA215002 Immunochemistry

CA209XXX Biochemical Methods

CA263XXX Pharmaceuticals

IDENTIFIERS: Neisseria meningitidis capsular polysaccharide purifn solubilization protein conjugate vaccine

DESCRIPTORS:

Immunostimulants...

adjuvants; purification of Neisseria meningitidis capsular polysaccharide for use in combination vaccines

Polysaccharides, biological studies...

capsular; purification of Neisseria meningitidis capsular polysaccharide for use in combination vaccines

Drug delivery systems...

carriers; purification of Neisseria meningitidis capsular polysaccharide for use in combination vaccines

Proteins... conjugates; purification of Neisseria meningitidis capsular polysaccharide for use in combination vaccines

Toxoids...

diphtheria, CRM197; purification of Neisseria meningitidis capsular polysaccharide for use in combination vaccines

Drug delivery systems...

freeze-dried; purification of Neisseria meningitidis capsular polysaccharide for use in combination vaccines

Neisseria meningitidis...

group A; purification of Neisseria meningitidis capsular polysaccharide for use in combination vaccines

Neisseria meningitidis...

group B; purification of Neisseria meningitidis capsular polysaccharide for use in combination vaccines

Neisseria meningitidis...

group C; purification of Neisseria meningitidis capsular polysaccharide for use in combination vaccines

Neisseria meningitidis...

group W-135, purification of Neisseria meningitidis capsular polysaccharide for use in combination vaccines

Neisseria meningitidis...

group Y; purification of Neisseria meningitidis capsular polysaccharide for use in combination vaccines

Buffers...

histidine; purification of Neisseria meningitidis capsular polysaccharide for use in combination vaccines Drug delivery systems...

ligs.; purification of Neisseria meningitidis capsular polysaccharide for use in combination vaccines

Carbohydrates, biological studies...

MenA and MenC; purification of Neisseria meningitidis capsular polysaccharide for use in combination vaccines

Physiological saline solutions...

phosphate-buffered; purification of Neisseria meningitidis capsular polysaccharide for use in combination vaccines

Bacteria (Eubacteria)... Solubilization... Precipitation (chemical)...

Neisseria meningitidis... Alcohols, biological studies... Ultrafiltration...

Haemophilus influenzae... Streptococcus pneumoniae... Hydrolysis...
Oligosaccharides, biological studies... Carriers... Biochemical molecules...

Antigens... Vaccines... Solvents...
purification of Neisseria meningitidis capsular polysaccharide for use in

purification of Neisseria meningitidis capsular polysaccharide for use : combination vaccines Filtration...

size; purification of Neisseria meningitidis capsular polysaccharide for use in combination vaccines
CAS REGISTRY NUMBERS:

7440-44-0 biological studies, activated; purification of Neisseria meningitidis capsular polysaccharide for use in combination vaccines

71-00-1 biological studies, buffer; purification of Neisseria meningitidis capsular polysaccharide for use in combination vaccines

64-17-5 21645-51-2 7732-18-5 biological studies, purification of Neisseria meningitidis capsular polysaccharide for use in combination vaccines 57-09-0 7784-30-7 purification of Neisseria meningitidis capsular

polysaccharide for use in combination vaccines

poryodecinariae for doe in comprinction vaccin

10/7/47 (Item 26 from file: 399) DIALOG(R)File 399:CA SEARCH(R)

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137045615 CA: 137(4)45615t JOURNAL

Isolation of oligosaccharides from a partial-acid hydrolysate of pneumococcal type 3 polysaccharide for use in conjugate vaccines

AUTHOR(S): Lefeber, Dirk J.; Gutierrez Gallego, Ricardo; Grun, Christian H.; Proietti, Daniela; D'Ascenzi, Sandro; Costantino, Paolo; Kamerling, Johannis P.; Vliegenthart, Johannes F. G.

LOCATION: Department of Bio-Organic Chemistry, Utrecht University, Bijvoet Center, 3508 TB, Utrecht, Neth.

JOURNAL: Carbohydr. Res. (Carbohydrate Research) DATE: 2002 VOLUME: 337 NUMBER: 9 PAGES: 819-825 CODEN: CRBRAT ISSN: 0008-6215 PUBLISHER ITEM IDENTIFIER: 0008-6215(02)00059-9 LANGUAGE: English PUBLISHER: Elsevier Science Ltd.

SECTION:

CA209XXX Biochemical Methods
CA233XXX Carbohydrates
IDENTIFIERS: Streptococcus carbohydrate oligosaccharide isolation

DESCRIPTORS:

Hydrolysis...

conjugate vaccine

acid; oligosaccharide isolation from a partial-acid hydrolyzate of pneumococcal type 3 polysaccharide for use in neoglycoprotein vaccines Polysaccharides, biological studies...

capsular; oligosaccharide isolation from a partial-acid hydrolyzate of pneumococcal type 3 polysaccharide for use in neoglycoprotein vaccines Glycoproteins...

anycoproteins; oligosaccharide isolation from a partial-acid hydrolyzate of pneumococcal type 3 polysaccharide for use in neoglycoprotein vaccines

Oligosaccharides, biological studies... Vaccines...

oligosaccharide isolation from a partial-acid hydrolyzate of pneumococcal type 3 polysaccharide for use in neoglycoprotein vaccines Ion exchange chromatography...

oligosaccharide isolation from a partial-acid hydrolyzate of pneumococcal type 3 polysaccharide fractionated by

Anion exchange chromatography... pH...

oligosaccharide isolation from a partial-acid hydrolyzate of pneumococcal type 3 polysaccharide purified by

Amperometry...
pulsed; oligosaccharide isolation from a partial-acid hydrolyzate of
pneumococcal type 3 polysaccharide purified by

Streptococcus pneumoniae...

type 3; oligosaccharide isolation from a partial-acid hydrolyzate of pneumococcal type 3 polysaccharide for use in neoglycoprotein vaccines CAS REGISTRY NUMBERS:

158129-72-7 oligosaccharide isolation from a partial-acid hydrolyzate of pneumococcal type 3 polysaccharide purified by

184867-28-5 repeating unit; oligosaccharides containing one to seven repeating units of

10/7/48 (Item 27 from file: 399)

DIALOG(R)File 399:CA SEARCH(R)
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132034363 CA: 132(4)34363e JOURNAI

Size determination of bacterial capsular oligosaccharides used to prepare conjugate vaccines

AUTHOR(S): Ravenscroft, Neil; Averani, Giovanni; Bartoloni, Antonella; Berti, Stefania; Bigio, Massimo; Carinci, Valeria; Costantino, Paolo; D'Ascenzi, Sandro; Giannozzi, Aldo; Norelli, Francesco; Pennatini, Carlo; Proietti, Daniela; Ceccarini, Costante; Cescutti, Paola

LOCATION: Chiron Vaccines SpA, I-53100, Siena, Italy

JOURNAL: Vaccine DATE: 1999 VOLUME: 17 NUMBER: 22 PAGES: 2802-2816 CODEN: VACCDE ISSN: 0264-410X PUBLISHER ITEM IDENTIFIER:

0264-410X(99)00092-4 LANGUAGE: English PUBLISHER: Elsevier Science Ltd. SECTION: CA215001 Immunochemistry IDENTIFIERS: bacteria capsule oligosaccharide size detn colorimetric assav vaccine DESCRIPTORS: Polysaccharides, biological studies... capsular; colorimetric assays for determination of the size of bacterial capsular oligosaccharides used to prepare conjugate vaccines Bacteria (Eubacteria) ... Neisseria meningitidis... Oligosaccharides, biological studies... Vaccines... colorimetric assays for determination of the size of bacterial capsular oligosaccharides used to prepare conjugate vaccines Haemophilus influenzae... type b; colorimetric assays for determination of the size of bacterial capsular oligosaccharides used to prepare conjugate vaccines 10/7/49 (Item 28 from file: 399) DIALOG(R)File 399:CA SEARCH(R) (c) 2011 American Chemical Society. All rts. reserv. 131078310 CA: 131(6)78310u JOURNAL Size fractionation of bacterial capsular polysaccharides for their use in conjugate vaccines AUTHOR(S): Costantino, Paolo; Norelli, Francesco; Giannozzi, Aldo; D'Ascenzi, Sandro; Bartoloni, Antonella; Kaur, Surinder; Tang, Dazhi; Seid, Robert; Viti, Stefano; Paffetti, Roberto; Bigio, Massimo; Pennatini, Carlo; Averani, Giovanni; Guarnieri, Valentina; Gallo, Eugenia; Ravenscroft, Neil; Lazzeroni, Carla; Rappuoli, Rino; Ceccarini, Costante LOCATION: Chiron Vaccines SpA, 53100, Siena, Italy JOURNAL: Vaccine DATE: 1999 VOLUME: 17 NUMBER: 9-10 PAGES: 1251-1263 CODEN: VACCDE ISSN: 0264-410X PUBLISHER ITEM IDENTIFIER: 0264-410X(98)00348-X LANGUAGE: English PUBLISHER: Elsevier Science Ltd. SECTION: CA263006 Pharmaceuticals CA215XXX Immunochemistry IDENTIFIERS: bacteria capsule polysaccharide antigen fractionation vaccine DESCRIPTORS: Vaccines... conjugate; size fractionation of bacterial capsular polysaccharides for use in conjugate vaccines Neisseria meningitidis... groups A and C; size fractionation of bacterial capsular polysaccharides for use in conjugate vaccines Capsule(microbial) ... Polysaccharides, biological studies ... size fractionation of bacterial capsular polysaccharides for use in conjugate vaccines Haemophilus influenzae... type b; size fractionation of bacterial capsular polysaccharides for use in conjugate vaccines 10/7/50 (Item 29 from file: 399) DIALOG(R)File 399:CA SEARCH(R) (c) 2011 American Chemical Society. All rts. reserv. 130050998 CA: 130(5)50998e JOURNAL A competitive enzyme-linked immunosorbent assay for measuring the levels of serum antibody to Haemophilus influenzae type b AUTHOR(S): Mariani, Massimo; Luzzi, Enrico; Proietti, Daniela; Mancianti,

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Silvia; Casini, Daniele; Costantino, Paolo; Van Gageldonk, Pieter; Berbers,
  LOCATION: Laboratorio di Immunochimica e Sierologia Sperimentale,
Dipartimento Immunologia, Centro Ricerche, I-53100, Siena, Italy
 JOURNAL: Clin. Diagn. Laboratory Immunol. DATE: 1998 VOLUME: 5 NUMBER: 5
  PAGES: 667-674 CODEN: CDIMEN ISSN: 1071-412X LANGUAGE: English
  PUBLISHER: American Society for Microbiology
  SECTION:
   CA215001 Immunochemistry
  IDENTIFIERS: ELISA antibody Haemophilus capsular polysaccharide
  DESCRIPTORS:
Blood analysis... Capsular polysaccharides...
    competitive ELISA for measuring human serum antibody to Haemophilus
    influenzae type b capsular polysaccharide
Vaccines..
    competitive ELISA for measuring human serum antibody to Haemophilus
    influenzae type b capsular polysaccharide in relation to
Antibodies... ELISA(immunosorbent assay)...
    competitive ELISA for measuring serum antibody to Haemophilus
    influenzae type b capsular polysaccharide
Serum albumin...
    conjugates, with Haemophilus capsular polysaccharide; for competitive
    ELISA measuring human serum antibody to Haemophilus influenzae type b
Capsular polysaccharides ...
    conjugates, with human serum albumin; for competitive ELISA measuring
    human serum antibody to Haemophilus influenzae type b
Microtiter plates ...
    precoated; with HSA-polysaccharide conjugate for competitive ELISA
    measuring human serum antibody to Haemophilus influenzae type b
Haemophilus influenzae...
    type b; competitive ELISA for measuring human serum antibody to
    Haemophilus influenzae type b capsular polysaccharide
 10/7/51
            (Item 30 from file: 399)
DIALOG(R)File 399:CA SEARCH(R)
(c) 2011 American Chemical Society. All rts. reserv.
              CA: 125(5)56224t
                                   PATENT
  Combined meningitis vaccine
  INVENTOR (AUTHOR): Ceccarini, Costante; Costantino, Paolo; D'Ascenzi,
Sandro: Norelli, Francesco: Giannozzi, Aldo
  LOCATION: Italy
  ASSIGNEE: Biocine S.P.A.
  PATENT: PCT International; WO 9614086 Al DATE: 960517
  APPLICATION: WO 951B1006 (951102) *GB 9422096 (941102)
  PAGES: 32 pp. CODEN: PIXXD2 LANGUAGE: English
  PATENT CLASSIFICATIONS:
    CLASS: A61K-039/02A; A61K-039/095B; A61K-039/102B
  DESIGNATED COUNTRIES: CA; JP; US DESIGNATED REGIONAL: AT; BE; CH; DE; DK
; ES; FR; GB; GR; IE; IT; LU; MC; NL; PT; SE
  SECTION:
    CA215002 Immunochemistry
  IDENTIFIERS: meningitis vaccine Hib MenC oligosaccharide conjugate, MenB
    oligosaccharide conjugate meningitis vaccine, DTP vaccine priming
   meningitis Hib MenC
  DESCRIPTORS:
Haemophilus influenzae, type b... Meningitis... Vaccines...
    combined vaccine for bacterial meningitis comprises Hib, MenC and MenB
    oligosaccharide conjugates
```

DTP vaccine; combined vaccine for bacterial meningitis comprises Hib,

Diphtheria... Tetanus... Whooping cough...

MenC and MenB oligosaccharide conjugates

Oligosaccharides, complexes...

of Haemophilus influenzae type B and Neisseria meningitidis serotype B and C; combined vaccine for bacterial meningitis comprises Hib, MenC and MenB oliopsaccharide conjugates

Neisseria meningitidis...

serotype B and C; combined vaccine for bacterial meningitis comprises Hib, MenC and MenB oligosaccharide conjugates

10/7/52 (Item 31 from file: 399)

DIALOG(R)File 399:CA SEARCH(R)

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123065627 CA: 123(6)65627n JOURNAL

Immunogenicity of meningococcal B polysaccharide conjugated to tetanus toxoid or CRM197 via adipic acid dihydrazide

AUTHOR(S): Bartoloni, Antonella; Norelli, Francesco; Ceccarini, Costante; Rappuoli, Rino; Costantino, Paolo

LOCATION: Biocine Research Center, IRIS, 53100, Siena, Italy JOURNAL: Vaccine DATE: 1995 VOLUME: 13 NUMBER: 5 PAGES: 463-70 CODEN: VACCDE ISSN: 0264-410X LANGUAGE: English SECTION:

CA263003 Pharmaceuticals

CA215XXX Immunochemistry

IDENTIFIERS: vaccine Neisseria polysaccharide carrier protein DESCRIPTORS:

Toxoids, tetanus...

conjugates; immunogenicity of meningococcal B polysaccharide conjugated to tetanus toxoid or CRM197 via adipic acid dihydrazide

Neisseria meningitidis,group B... Polysaccharides,conjugates,biological studies... Proteins,specific or class, CRM 197, conjugates... Toxins,diphtheria... Vaccines...

immunogenicity of meningococcal B polysaccharide conjugated to tetanus toxoid or CRM197 via adipic acid dihydrazide

CAS REGISTRY NUMBERS:

1071-93-8 immunogenicity of meningococcal B polysaccharide conjugated to tetanus toxoid or CRM197 via adipic acid dihydrazide

10/7/53 (Item 32 from file: 399)

DIALOG(R)File 399:CA SEARCH(R)

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119269016 CA: 119(25)269016g PATENT

Conjugates formed from heat-shock proteins and oligo- or polysaccharides for vaccine against bacterial infection

INVENTOR (AUTHOR): Rappuoli, Rino; Costantino, Paolo; Viti, Stefano; Norelli, Francesco

LOCATION: Italy

ASSIGNEE: Biocine Sclavo SPA

PATENT: PCT International ; WO 9317712 A2 DATE: 930916

APPLICATION: WO 93EP516 (930308) *IT 92F158 (920306)

PAGES: 69 pp. CODEN: PIXXD2 LANGUAGE: English

PATENT CLASSIFICATIONS:

CLASS: A61K-047/48A; C07K-015/00B

DESIGNATED COUNTRIES: AT; AU; BB; BG; BR; CA; CH; CZ; DE; DK; ES; FI; GB; HU; JP; KP; KR; LK; LU; MG; MN; MN; NL; NO; NZ; PL; PT; RO; RU; SD; SE; SK; UA; US DESIGNATED REGIONAL: AT; BE; CH; DE; DK; ES; FR; GB; GR; IE; IT; LU; MC; NL; PT; SE; BF; BJ; CF; CG; CI; CM; GA; GN; ML; MR; SN; TD; TG SECTION:

CA215002 Immunochemistry

```
IDENTIFIERS: heat shock protein oligosaccharide conjugate vaccine,
    polysaccharide heat shock protein conjugate vaccine, Helicobacter heat
    shock protein DNA cloning, sequence Helicobacter heat shock protein
  DESCRIPTORS:
Vaccines...
    against bacterial infection, conjugates of heat-shock proteins with
    oligo- or polysaccharides for
Deoxyribonucleic acid sequences... Protein sequences...
    for heat-shock protein of Helicobacter pylori
Campylobacter pyloridis ...
    heat-shock protein of, conjugates with oligo- or polysaccharides, for
    vaccines against bacterial infection
Proteins, specific or class, heat-shock...
```

hspR65, conjugates with oligo- or polysaccharides, for vaccines against bacterial infection

Proteins, specific or class, heat-shock...

hspR70, conjugates with oligo- or polysaccharides, for vaccines against bacterial infection

Neisseria meningitidis, group C...

oligosaccharides of, conjugates with heat-shock proteins, for vaccines against bacterial infection

Plasmid and Episome..

pHp60G2 and pHp60G5, heat-shock protein of Helicobacter pylori in relation to

Oligosaccharides, conjugates... Polysaccharides, conjugates, compounds... with heat-shock proteins, for vaccines against bacterial infection Proteins, specific or class, heat-shock, complexes...

with oligo- or polysaccharides, for vaccines against bacterial infection

CAS REGISTRY NUMBERS:

10/7/54

SECTION:

151441-76-8 amino acid sequence of and cloning of DNA for, oligo- or polysaccharide conjugates with heat-shock proteins for vaccines in relation to

9026-28-2D heat-shock protein fusion products, of phage MS2, with Helicobacter pylori heat-shock protein

151243-14-0 nucleotide sequence of and cloning of, oligo- or polysaccharide conjugates with heat-shock proteins for vaccines in relation to

(Item 33 from file: 399) DIALOG(R)File 399:CA SEARCH(R) (c) 2011 American Chemical Society. All rts. reserv.

CA: 117(21)210275d JOURNAL

Mycobacterial heat-shock proteins as carrier molecules. II: The use of the 70-kDa mycobacterial heat-shock protein as carrier for conjugated vaccines can circumvent the need for adjuvants and Bacillus Calmette Guerin priming

AUTHOR(S): Barrios, Christy; Lussow, Alexander R.; Van Embden, Jan; Van der Zee, Ruurd; Rappuoli, Rino; Costantino, Paolo; Louis, Jacques A.; Lambert, Paul Henri; Del Giudice, Giuseppe

LOCATION: World Health Organ. Immunol. Res. Training Cent., University Geneva, CH-1211, Geneva, Switz.

JOURNAL: Eur. J. Immunol. DATE: 1992 VOLUME: 22 NUMBER: 6 PAGES: 1365-72 CODEN: EJIMAF ISSN: 0014-2980 LANGUAGE: English

CA215002 Immunochemistry

IDENTIFIERS: Mycobacterium heat shock protein vaccine carrier DESCRIPTORS:

Immunostimulants, adjuvants...

antibody formation to vaccines conjugated to mycobacterial heat-shock

```
protein carrier after immunization in relation to
Polysaccharides, biological studies...
    capsular, of Neisseria meningitidis, mycobacterial heat-shock protein
    conjugates immunization with, antibody formation induction by, immune
    adjuvants and BCG priming in relation to
Plasmodium falciparum...
    circumsporozoite antigen of, peptide from, mycobacterial heat-shock
    protein conjugates, immunization with, antibody formation after, immune
    adjuvants and BCG priming in relation to
Proteins, specific or class, hsp 70...
    conjugates, mycobacterial, with vaccines, antibodies to, formation of,
    immune adjuvants and BCG priming in relation to
Mycobacterium tuberculosis...
    heat-shock protein 70 from, as immunostimulatory vaccine carrier
Lymphocyte, T-cell...
    in antibody formation to vaccines conjugated to mycobacterial
    heat-shock protein carrier
Vaccines...
    oligosaccharides and peptide conjugates with mycobacterial heat-shock
    protein carrier in relation to
Neisseria meningitidis...
    oligosaccharides from, mycobacterial heat-shock protein conjugates,
    immunization with, antibody formation induction by, immune adjuvants
    and BCG priming in relation to
Proteins, specific or class, hsp 65...
    preimmunization with, antibody formation to vaccines conjugated to
    mycobacterial heat-shock protein carrier enhancement by
Antigens, CS (circumsporozoite)...
    synthetic peptide from, mycobacterial heat-shock protein conjugates,
    immunization with, antibody formation induction by, immune adjuvants
    and BCG priming in relation to
Antibodies...
    to vaccines conjugated to mycobacterial heat-shock protein, formation
    of, immune adjuvants and BCG priming in relation to
Oligosaccharides, conjugates... Peptides, conjugates, compounds...
    with heat-shock protein 70 from Mycobacterium, immunostimulation by,
    vaccines in relation to
  CAS REGISTRY NUMBERS:
143180-71-6D mycobacterial heat-shock protein conjugates, immunization
    with, antibody formation induction by, immune adjuvants and BCG priming
    in relation to
 10/7/55
             (Item 34 from file: 399)
DIALOG(R)File 399:CA SEARCH(R)
```

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117189760 CA: 117(19)189760w JOURNAL

Development and phase 1 clinical testing of a conjugate vaccine against meningococcus A and C

AUTHOR(S): Costantino, Paolo; Viti, Stefano; Podda, Audino; Velmonte, Melecia Antonio; Nencioni, Luciano; Rappuoli, Rino

LOCATION: Sclavo Res. Dev. Vaccines, 53100, Siena, Italy JOURNAL: Vaccine DATE: 1992 VOLUME: 10 NUMBER: 10 PAGES: 691-8 CODEN: VACCDE ISSN: 0264-410X LANGUAGE: English SECTION:

CA215002 Immunochemistry CA263XXX Pharmaceuticals IDENTIFIERS: meningococcus vaccine DESCRIPTORS:

Vaccines...

to meningococcus

with oligosaccharidic haptens, preparation of, as vaccine against gram-pos.

10/7/57 (Item 36 from file: 399) DIALOG(R)File 399:CA SEARCH(R) (c) 2011 American Chemical Society. All rts. reserv.

104205102 CA: 104(23)205102x JOURNAL

Proteins, specific or class, CRM 197, conjugates...

and gram-neg. bacterial infection

glycoprotein antigens as

A molecular model of artificial glycoprotein with predetermined multiple immunodeterminants for gram-positive and gram-negative encapsulated bacteria

AUTHOR(S): Porro, Massimo; Costantino, Paolo; Giovannoni, Franco; Pellegrini, Vittoria; Tagliaferri, Lucia; Vannozzi, Francesca; Viti,

LOCATION: Bact. Vaccine Dep., Sclavo SpA, 53100, Siena, Italy JOURNAL: Mol. Immunol. DATE: 1986 VOLUME: 23 NUMBER: 4 PAGES: 385-91 CODEN: MOIMD5 ISSN: 0161-5890 LANGUAGE: English

SECTION: CA215002 Immunochemistry

CA210XXX MICROBIAL, ALGAL, AND FUNGAL BIOCHEMISTRY

IDENTIFIERS: oligosaccharide protein conjugate bacteria multivalent immunogenicity DESCRIPTORS:

Antigens... Glycoproteins...

bacterial oligosaccharide-protein CRM 197 conjugate as synthetic, multivalent immunogenicity of, vaccines in relation to

Vaccines...

bacterial oligosaccharide-protein CRM 197 conjugates as, multivalent determinants of

Proteins, CRM 197...

conjugate with Neisseria meningitidis and Streptococcus pneumoniae oligosaccharides, multivalent immunogenicity of, vaccines in relation

Oligosaccharides...

of Neisseria meningitidis and Streptococcus pneumoniae, conjugates with protein CRM 197, multivalent immunogenicity of, vaccines in relation to Capsule, microbial...

of Neisseria meningitidis and Streptococcus pneumoniae,

oligosaccharides of, conjugates with protein CRM 197, multivalent immunogenicity of, vaccines in relation to

Streptococcus pneumoniae...

oligosaccharides of Neisseria meningitidis and, conjugates with protein CRM 197, multivalent immunogenicity of, vaccines in relation to Neisseria menincitidis...

oligosaccharides of Streptococcus pneumoniae and, conjugates with

Items Description

protein CRM 197, multivalent immunogenicity of, vaccines in relation to Toxins, diphtheria...

protein CRM 197 related to, bacterial oligosaccharides conjugates with, multivalent immunogenicity of, vaccines in relation to

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                    S8 AND (CAPSULAR OR POLYSACCHARIDE OR NEISSERIA OR M-
                ENINGITIDIS)
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    $2.64 Estimated cost File144
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